

Effect of Gum-Chewing after Cesarean Section on Gastrointestinal Function Recovery: A Systematic Review and Meta-Analysis of **Randomized Trials**

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How to cite this paper: Zhang, L., Heng, Y.L., Hu, H. and Liang, X. (2019) Effect of Gum-Chewing after Cesarean Section on Gastrointestinal Function Recovery: A Systematic Review and Meta-Analysis of Randomized Trials. Yangtze Medicine, 3, 79-89

https://doi.org/10.4236/ym.2019.32009

Received: October 21, 2018 Accepted: March 19, 2019 Published: March 22, 2019

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Abstract

Objective: To evaluate the effect of chewing sugar-free gum on gastrointestinal function recovery after cesarean section. Methods: Randomized controlled trials comparing the use of chewing gum in postoperative recovery with a control group were retrieved from the databases including Pubmed, Cochrane Library, Science Direct, Web of Science, CBM, CNKI, Wanfang and VIP, et al. After screening literatures, evaluating the quality of studies, extracting data, the RevMan5.3 software was used for data analysis. Results: A total of 13 RCTs including 2233 patients were analyzed in the study. The results showed that chewing gum after cesarean delivery can effectively shorten the recovery time of bowel sounds, passage of flatus and first defecation (all P < 0.05). Conclusion: Chewing sugar-free gum after cesarean section can promote the early recovery of gastrointestinal function, but the side effects of chewing gum are still unclear, which needs more clinical, large sample and high-quality studies to further verify.

Keywords

Cesarean Section, Gum-Chewing, Gastrointestinal Function, Randomized Controlled Trial, Systematic Review

1. Introduction

In recent years, with the development of medical technology and related support policies, the cesarean section rate in the world has tended to rise [1]. According to the survey conducted by WHO in 2010, the rate of cesarean section in China

is the highest among 9 countries in Asia, up to 46%. From 2008 to 2014, the total annual delivery rate of cesarean section in China increased by 34.9% [2]. However, cesarean section is prone to cause a series of postoperative complications: Postoperative ileus is an impaired condition of gastrointestinal, most likely to occur with day 4 postoperatively; the average incidence of postoperative ileus is about 20% [3]. A large number of studies have shown that early feeding can effectively promote the recovery of gastrointestinal function after cesarean section, but some patients performed high intolerance [4] [5], and delayed feeding is related to increase cell break-down, delay the time of wound healing and increase the risk of infection, thereby increasing medical costs [6] [7]. As early as 1992, Soffer pointed out that sham feeding could promote the secretion of saliva and pancreatic juice through the head-vagus reflex, thus promoting the early recovery of gastrointestinal function [8]. More studies have shown that chewing gum after abdominal surgery, including cesarean section, can effectively promote early recovery of gastrointestinal function and reduce postoperative complications [9]-[23]. At present, domestic meta-analysis is required more trials to provide sufficient evidence. The aim of this study was to evaluate the effect of chewing gum on the early recovery of gastrointestinal function after cesarean section by the method of evidence-based medicine, and provide a basis for further clinical exploration of the early recovery of gastrointestinal function in patients after cesarean section.

2. Materials and Methods

2.1. Inclusion Criteria

1) Study Design: randomized controlled trials (RCTs) about chewing gum in early stage after cesarean section were included in Chinese and English. 2) Participants: after cesarean section, the patients were in good health, conscious and had no serious complications (such as gestational hypertension, severe eclampsia, diabetes mellitus, etc.). 3) Interventions: the intervention group chewed sugar free gum after cesarean section, while the control group received routine nursing. 4) Outcome indicators (satisfying one of them): a) recovery time of bowel sounds; b) first time of passage of flatus; c) first time of defecation; d) first feeling of hunger.

2.2. Exclusion Criteria

The studies in which has one of the following conditions will be excluded, the interventions consisted of chewing gum in combination with other interventions; have no access to the original literature; incorrect use of data collection and statistical methods.

2.3. Search Strategy

We tried to plan, perform and report this meta-analysis in comply with PRISMA guideline [24]. Foreign language databases and Chinese databases were searched

from their inception until July 2018 for randomized controlled trials about chewing gum in early stage after cesarean section, included Pubmed, Cochrane Library, Web of Science, Science Direct, Chinese Biomedical Literature Database (CBM), China National Knowledge Infrastructure(CNKI), WanFang database and VIP database, etc. Searchterms: "cesarean section/delivery", "gum-chewing/chewing", "randomized controlled trial/RCT" and "gastrointestinal function/gastrointestinal complications", and Mesh keywords, free words and Boolean logic operator conjunctions are combined in accordance to the instructions of all kinds of database. Retrieval words of similar meaning, such as "cesarean section" and "cesarean delivery" are linked with "or", but "cesarean section" and "gum-chewing" are linked with "and". In addition, the references of the retrieval studies and previous reviews were reviewed, computer retrieval is the main method, supplemented by the secondary literature traceability method to expand the retrieval, so as to avoid missing the studies not captured by electronic searches.

2.4. Study Selection and Quality Evaluation

1) Study screening and data extraction: According to the consistent inclusion and exclusion criteria, the data was selected and extracted by two researchers independently, the two researchers have learned knowledge about evidence-based medicine, and received standardized training. Extract contents including basic information (author, year of publication, country, baseline situation, etc.), research design, sample size, interventions, outcome indicators, etc. Any disagreement was resolved by discussion with a third reviewer. 2) Quality assessment: the two researchers independently evaluated the included studies according to Cochrane Assessment Manual 5.1.0 Assessment Criteria. The evaluation items included: random sequence generation, allocation hiding, blind method, data integrity of outcome indicators, selective reporting of findings and other sources of bias. A: fully meet the above criteria, the risk of a variety of bias is low; B: partially meet the above criteria, bias risk is moderate; C: completely dissatisfied with the above criteria, bias risk is higher. Only "A" and "B" studies were included in this study.

2.5. Statistical Analysis

Meta-analysis was performed using the RevMan5.3 software, provided by Cochrane collaboration network. Weighted mean difference (MD) was used for measurement data, such as recovery time of bowel sounds, first time of passage of flatus, first time of defecation, first feeling of hunger. Relative Risk (RR) was used for enumeration data. 95% CI was calculated for all analyses. Firstly, Chi-square (X^2) was used to confirm whether there was heterogeneity among the studies. If there was no significant statistical heterogeneity (P > 0.1, $\hat{I} < 50\%$), the fixed effect model was adopted. If there was heterogeneity (P < 0.1, $\hat{I} \ge$ 50%), the random effect model was used to combine with the effect size. Descriptive analysis was adopted if heterogeneity was too obvious and can't be judged.

3. Results

3.1. Study Selection and Study Characteristics

A total of 443 trials about chewing-gum after cesarean section were initially retrieved, and 49 papers were included through preliminary reading literature and abstracts, removing summaries, case reports and small methods. Finally, 13 trials were assessed for eligibility after further reading. The flow of study identification is shown in **Figure 1**. **Table 1** shows the characteristics of the included studies. The intervention group chewed sugar-free gum after cesarean section, while the control group received routine nursing, *i.e.*, fluidized diet after a wake anesthesia or the first exhaust after cesarean section and gradually transited to general diet.

Table 1. Characteristics of the included studies	Table 1.	Characteristics	of the	included	studies.
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Included studies	Year	Nation	(Interv	le size vention trol)		terventions start Chewing method	Control group	Outcome indicator
Liang [11]	2007	China	60	60	From delivery	Every 2 h 15 min each, 3 times/d	routine	23
Abd-EI-Maeboud [12]	2009	Egypt	93	107	From 2 h postoperatively	2 h/every times 15 min each, until first flatus	routine	123
Luo [13]	2010	China	150	150	From 2h postoperatively	4 times/d, 10 - 15 min each Until 3 days after CD	routine	123
Wang [14]	2011	China	116	117	From 2 h postoperatively	Every 2 h, 0 - 15 min each until first flatus	routine	12
Ledari [15]	2012	Iran	50	50	From 6 h postoperatively	3 times/d At least 1 h each Until discharge	routine	1234
Ledari [16]	2013	Iran	30	30	From 6 h postoperatively	3 times/d At least 1 h each Until discharge	routine	1234
Rashad [17]	2013	Saudi Arabia	30	30	From posto-anesthesia	3 times/d 30 min each Until discharge	routine	123
Ajuzieogu [18]	2014	Nigeria	90	90	From the first day postoperatively	3 times/d 30 min each Until 5 days after CD	routine	123
Jian Tao [19]	2015	China	40	40	From 2 h postoperatively	Every 2 h, 15 min each Until first flatus	routine	123
Lang [20]	2017	China	60	60	From 2 h postoperatively	3 times/d 30 min each Until first flatus	routine	23
Feng [21]	2017	China	130	130	From 15 min postoperatively	Every 30 min 10 - 15 min each Until first flatus	routine	123
Wang [22]	2018	China	110	110	From 2 h postoperatively	4 times/d 15 min each	routine	123
Magdy [23]	2018	Egypt	100	100	From 2 h postoperatively	Every 2 h, 10 min each Until first flatus	routine	123

①: recovery time of bowel sounds; ②: the time of first passage of flatus; ③: the time of first defection; ④: the time of first feeling of hunger.

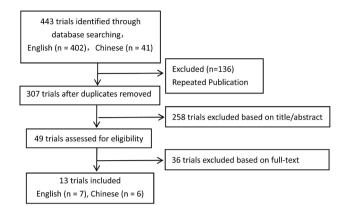


Figure 1. Flow diagram of studies identified in the systematic review.

3.2. Methodological Quality Evaluation of the Literature

Therefore 13 trials, including 2233 patients were analyzed, of which 7 trials were in English and 6 in Chinese. The methodological quality of the study was evaluated in **Table 2**. It showed the qualities of the 3 studies were grade A and 10 were grade B.

3.3. Results of Systematic Reviews

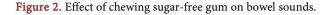
3.3.1. Effect of Chewing Sugar-Free Gum on Bowel Sounds

11 studies [10]-[17] [19] [20] [21] reported the effect of chewing gum post operatively on the recovery time of bowel sounds. 6 trials [10] [11] [12] [17] [20] [21] chewing gum within 2 h of delivery, and were analyzed quantitatively, the combined results showed that there was heterogeneity between the studies (P < 0.00001, $I^2 = 97\%$), so a randomized effect model was used for meta-analysis. The results showed that the recovery time of bowel sounds of postoperatively chewing gum was earlier than that of routine nursing group, and the difference was statistically significant [MD = -6.27, 95% CI (-7.98, -4.56), Z = 7.18, P < 0.00001], as shown in Figure 2. The sensitivity analysis showed the difference in each group had statistically significant after removing the study with the smallest sample size, indicating that the results were stable. And Ledari *et al.* [13] [14] [15] [16] [19] reported that chewing gum at different times after surgery the results also showed statistically significant.

3.3.2. Effect of Chewing Sugar-Free Gum on the Passage of Flatus

The effect of postoperatively chewing gum on anal exhaust time was reported in 13 trials, 7 studies [10] [11] [12] [17] [18] [20] [21] could be quantitatively analyzed. Heterogeneity was examined by Chi-square, found there was heterogeneity in each group (P < 0.00001, $\vec{F} = 99\%$), so the random effect model was adopted. The results showed that the first time of passage of flatus in chewing gum group was shorter than that in control group, and the difference was statistically significant [MD = -8.66, 95% CI (-14.12, -3.20), Z = 3.11, P = 0.002], as shown in **Figure 3**. At the same time, in Liang Junhua and other studies [9] [13] [14] [15] [16] [19], the results were different between the two groups (P < 0.05).

	Interver	ntion gr	oup	Cont	rol gro	oup		Mean Difference		Mea	n Differei	ice	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI		IV, Ra	indom, 9	5% CI	
Abd-Ei-Maeboud 2009	10.9	2.7	93	15.8	3.7	107	16.8%	-4.90 [-5.79, -4.01]					
Jian Tao 2015	6.9	1.7	40	12.8	2.5	40	16.7%	-5.90 [-6.84, -4.96]		-			
Luo 2010	6.67	1.4	150	13.14	2.3	150	17.4%	-6.47 [-6.90, -6.04]					
Magdy 2018	10	2.6	100	21.6	4.2	100	16.6%	-11.60 [-12.57, -10.63]					
Wang 2011	12.8	3.1	116	16.7	8.4	110	15.1%	-3.90 [-5.57, -2.23]		-			
Wang 2018	7.3	1.3	110	12	1.7	110	17.4%	-4.70 [-5.10, -4.30]					
Total (95% CI)			609			617	100.0%	-6.27 [-7.98, -4.56]		•			
Heterogeneity: Tau ² = 4.	34; Chi ² = 1	184.80,	df = 5 (l	< 0.000	001); l	² = 97%			-10				10
Test for overall effect: Z	= 7.18 (P <	0.0000	1)							-5 ervention gro	oup Con	5 trol group	



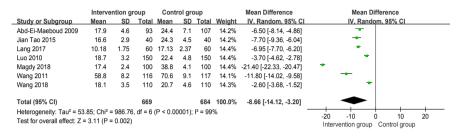


Figure 3. Effect of chewing sugar-free gum on the passage of flatus.

Table 2. Assessment of risk of bias.

Inclusion studies	Random sequence generation	Blinding of participants	outcome	Incomplete outcome data	Selective reporting	Other bias	Quality grade
Liang [11]	unclear	unclear	unclear	low	low	unclear	В
Abd-EI-Maeboud [12]	low	low	low	low	low	low	А
Luo [13]	unclear	unclear	unclear	low	low	unclear	В
Wang [14]	low	unclear	unclear	low	low	low	В
Ledari [15]	low	unclear	low	low	low	low	В
Ledari [16]	unclear	unclear	unclear	low	low	unclear	В
Rashad [17]	unclear	unclear	unclear	low	low	unclear	В
Ajuzieogu [18]	low	low	low	low	low	low	А
Jian Tao [19]	low	low	low	low	low	low	А
Lang [20]	low	unclear	unclear	low	low	low	В
Feng [21]	low	unclear	unclear	low	low	low	В
Wang [22]	low	unclear	unclear	low	low	low	В
Magdy [23]	low	unclear	unclear	low	low	low	В

A: Fully meet the above criteria, the risk of a variety of bias is low; B: Partially meet the above criteria, bias risk is moderate.

3.3.3. Effect of Chewing Gum on the Time of First Defecation

12 studies [9] [10] [11] [13]-[21] reported the effect of postoperatively chewing gum on the first time of defecation. 6 studies [10] [11] [17] [18] [20] [21] were analyzed quantitatively. Heterogeneity showed that random effect model should be selected to analyze (P < 0.00001, $\vec{I} = 99\%$). The results showed that the defecation time of the chewing sugar-free gum group was earlier than that of the routine nursing group, and the difference was statistically significant [MD =

-9.06, 95% *CI* (-15.59, -2.53), *Z* = 2.72, *P* = 0.007], as shown in **Figure 4**. In the meanwhile, in the studies of Liang Junhua *et al.* [9] [13] [14] [15] [16] [19], the difference was also statistically significant, indicating that early chewing gum can promote defecation time ahead.

3.3.4. Effect of Chewing Gum on the Time of First Feeling of Hunger

Two trials [13] [14] reported the effect of postoperatively chewing gum on the time of first feeling of hunger. There was no heterogeneity between two studies (P = 0.18, $\vec{l} = 43\%$), therefore, the fixed effect model was used. The results showed that chewing gum after cesarean section can promote patients feel hunger and then start early feeding [MD = -3.75, 95% CI(-5.99, -1.50), Z = 3.27, P = 0.001], as shown in Figure 5.

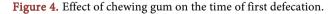
4. Discussion

4.1. Methodological Quality of the Included Studies

13 trials were included in this study, the quality of the included trials was medium. 9 trials [12] [14] [15] [18]-[23] produced random sequences by computer-ganerated random sequence. 4 trials [11] [13] [16] [17] just mentioned random allocation, how random sequence generated was unclear, because it was difficult to achieve double-blind for intervention group and control group, three studies [12] [18] [19] conducted a single-blind. One trial [15] reported the missing rate and missing reasons. The baseline datas of age, gestational age, education level and parity were compared between the intervention group and the control group. The results showed that the baseline datas were comparable between the two groups (P > 0.05).

However, the clinical heterogeneity between these studies was large, and the specific program of chewing gum was obviously different. Most studies began chewing gum within 2 h of cesarean delivery, while a few studies [15] [16] [17] [18] began after 6 h or more of surgery, whether it made sense in early stage was unclear. Xie Minyi *et al.*, [25] investigated the effect of chewing gum at different

	Interver	ntion gr	oup	Cont	rol gro	bup		Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
Abd-Ei-Maeboud 2009	21.1	4.7	93	30	8.2	107	20.0%	-8.90 [-10.72, -7.08]	-
Jian Tao 2015	42.2	5.6	40	43.4	0.9	40	20.0%	-1.20 [-2.96, 0.56]	
Luo 2010	54.2	6.9	150	66.4	8.1	150	20.0%	-12.20 [-13.90, -10.50]	+
Magdy 2018	28.1	4.7	100	49.4	6.2	100	20.1%	-21.30 [-22.82, -19.78]	+
Wang 2018	48.7	6	110	57.7	6.8	110	20.0%	-9.00 [-10.69, -7.31]	-
Total (95% CI)			493			507	100.0%	-10.53 [-17.17, -3.89]	•
Heterogeneity: Tau ² = 56			, df = 4	(P < 0.00	0001);	l² = 99	%	-	-20 -10 0 10 20
Test for overall effect: Z =	= 3.11 (P =	0.002)							Intervention group Control group



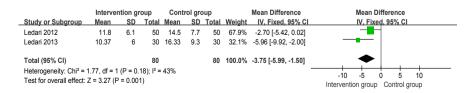


Figure 5. Effect of chewing gum on the time of first feeling of hunger.

time windows on the recovery of gastrointestinal function after gastrointestinal surgery, the results showed that the level of serum gastrin in group A (chewing gum at postoperative 2 h) were higher than those in group B (chewing gum at postoperative 6 h) and the incidence of abdominal distention was lower in group A than that in group B (P < 0.05), indicated that chewing gum the earlier, the effect is better. Some studies [15] [16] [17] [18] stopped chewing gum after discharge or 5 days of delivery, which was different from the general study, and it was not clear whether chewing gum was still meaningful after patients returned to normal diet, while in some studies [15] [16], patients chewed gum for more than one hour, the time of chewing was too long and the frequency was too much can easily lead to masseter muscle fatigue, abdominal distension, dizziness, nausea and other adverse reaction.

4.2. Effect of Chewing Gum on Gastrointestinal Function Recovery after Cesarean Section

The gastrointestinal adverse reactions such as nausea, vomiting and abdominal distention are more likely to occur during the period from inhibition to full recovery of gastrointestinal function after cesarean section. Recovering gastrointestinal function as soon as possible is of great significance in preventing gastrointestinal complications after cesarean section and promoting the health of maternal and infant. Chewing gum, as a non pharmacological methods is convenient and safe, it not only doesn't increase the burden of gastrointestinal tract and cause a series of complications, but also stimulate gastrointestinal motility, which is easily accepted by patients. Studies [14] [26] have showed that chewing gum can significantly increase the levels of gastrin and motilin in peripheral gastrointestinal hormones on the first and second day after operation, thereby promoting gastrointestinal motility. And, the majority of cesarean section objects are young mothers with good digestive function, the way of anesthesia is epidural anesthesia, postoperative patients have clear consciousness, and the possibility of vomiting, asphyxia and other complications caused by eating are low. It indicates that maternal have the physiological basis of early feeding, it's safe enough for maternal to chewing gum after delivery [27].

Related meta-analysis [28] [29] showed that chewing gum could promote the early recovery of gastrointestinal function after cesarean section, but the start time and frequency of chewing gum were different. In most of the included studies, chewing gum was given within 2 h of delivery, at least 3 times one day, 15 - 30 minutes each time, and until the first bowel sounds was heard. Quantitative analysis of the combined studies was carried out. Otherwise, descriptive analysis was selected. The results showed that chewing gum could promote the early recovery of maternal gastrointestinal function. It was consistent with the results of review by Perriera *et al.* [30]. And, Liang Junhua *et al.*, [11] [21] [22] showed that chewing gum postoperative can effectively reduce the incidence of nausea, vomiting, abdominal distention and other adverse gastrointestinal reactions. Abd-EI-Maeboud *et al.*, [12] [20] [21] showed that chewing gum after surgery

can shorten hospitalization time. However, study [18] showed that it has no significant difference in hospitalization time compared with control group. Study [31] showed that mastication activates histamine nervous system conduction signals and promotes histamine production, thereby reducing hunger by inhibiting food intake by receptors located in the center of satiety. Whether the feeling of hunger can be used as a criterion for evaluating gastrointestinal functional recovery needs to be further discussed.

4.3. Limitations of This Systematic Review

Because this study only retrieves the published Chinese and English trials, some trials may not be included. There are great differences in the sample and methodology of the included studies, and the intervention schemes are not consistent, which may have a certain impact on the results of the study. Sugar-free gums containing artificial sweetener sorbitol and other alcohols may have side effects such as bloating, flatulence and abdominal colic [32]. However, in the 13 trials, there was no study reported the side effect of chewing gum, the effect of chewing gum on the recovery of gastrointestinal function remains to be further explored.

5. Conclusion

Chewing gum as a simple and convenient intervention can effectively promote the recovery of bowel sounds after cesarean section, and shorten the first time of passage of flatus and the defecation. In the future, we can expand the sample size and carry out high-quality randomized controlled trials to validate the effect of standardized chewing gum intervention program (start time, frequency, chewing dose, duration, etc.) on promoting the recovery of maternal gastrointestinal function.

Conflicts of Interest

The authors declare no conflicts of interest regarding the publication of this paper.

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