# Review: Smooth-muscle relaxants improve symptoms and reduce pain in the irritable bowel syndrome

Poynard T, Regimbeau C, Benhamou Y. Meta-analysis of smooth muscle relaxants in the treatment of irritable bowel syndrome. Aliment Pharmacol Ther. 2001 Feb;15:355-61.

## QUESTION

In patients with the irritable bowel syndrome (IBS), are smooth-muscle relaxants safe and efficacious for global improvement of symptoms?

## DATA SOURCES

Randomized controlled trials were identified by searching MEDLINE with the terms colonic disease, functional, irritable bowel syndrome, and randomized trial; *Current Contents*; and bibliographies of relevant trials, meta-analyses, and reviews. Drug companies were also contacted.

## STUDY SELECTION

Trials were selected if they were double-blind studies that compared muscle relaxants with placebo;  $\geq 51\%$  of patients had IBS; the drugs studied had myorelaxant properties; and  $\geq 1$  end point of global assessment of symptoms, abdominal pain, constipation, or abdominal distention was included.

# DATA EXTRACTION

Data were extracted on study quality, patient characteristics and numbers, drug and dosage, treatment duration, and outcomes.

# MAIN RESULTS

23 trials (1888 patients) met the inclusion criteria. 6 drugs were studied: cimetropium

bromide (5 trials), hyoscine butyl bromide (3 trials), mebeverine (5 trials), otilium bromide (4 trials), pinaverium bromide (2 trials), and trimebutine (4 trials). All drugs analyzed together and each drug analyzed separately, except pinaverium alone, showed improvement in global symptom ratings (Table). All drugs combined (and otilium and cimetropium alone) reduced pain (Table). Abdominal distention was improved with all drugs (Table), but constipation or transit was not. The percentage of patients with adverse

effects did not differ for all drugs compared with placebo or for individual drugs.

## CONCLUSION

Smooth-muscle relaxants improve the global rating of symptoms and reduce pain in patients with the irritable bowel syndrome.

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# Smooth-muscle relaxants vs placebo for the irritable bowel syndrome (trial duration 3 d to 24 wk)\*

Improvements	Drug	Number of studies	Unweighted event rates		RBI (95% CI)	NNT (CI)
			Drug	Placebo		
Global	All	22	56%	38%	49% (37 to 61)	6 (5 to 8)
	Mebeverine	4	52%	37%	47% (9 to 84)	4 (6 to 31)
	Cimetropium	5	60%	40%	63% (29 to 92)	4 (3 to 9)
	Trimebutine	4	53%	24%	98% (63 to 117)	5 (4 to 7)
	Otilonium	3	97%	34%	60% (33 to 87)	5 (4 to 9)
	Hyoscine	3	53%	42%	26% (7 to 124)	10 (2 to 32)
Pain	All	11	53%	41%	30% (16 to 45)	9 (6 to 16)
	Otilium	3	47%	29%	62% (30 to 96)	7 (4 to 12)
	Cimetropium	1	87%	27%	198% (81 to 253)	2 (2 to 5)
Distention	All	6	44%	35%	26% (6 to 46)	12 (7 to 46)

<sup>\*</sup>Abbreviations defined in Glossary; RBI, NNT, and CI calculated from data in article.

# COMMENTARY

IBS represents a heterogeneous group of disorders presenting with similar clinical symptoms. Altered motility is thought to be important in a subset of patients with IBS, and this concept forms the foundation for the use of smooth-muscle relaxants.

3 recent meta-analyses, including the review by Poynard and colleagues, have evaluated smooth-muscle relaxants for IBS (1, 2). Each found such agents better than placebo for global improvement and abdominal pain. The latest, the review by Poynard and colleagues, differs from its predecessors by excluding peppermint oil and dyciclomide bromide and including 6 additional randomized, non—English-language trials. Although smooth-muscle relaxants appear superior to placebo, Poynard and colleagues suggest that only approximately 50% of treated patients will experience global improvement and only 40% will have an improvement in abdominal pain. In addition, the results do not indicate which individuals are most likely to improve with these agents. Perhaps most important, only 1 of the drugs included in this meta-analysis, hyoscine, is

available in the United States, and it showed the least benefit for global improvement or abdominal pain over placebo.

If one accepts the issues noted above, it seems reasonable to conclude that as a class, smooth-muscle relaxants can benefit a subset of patients with IBS. Unfortunately, whether the results are generalizable to agents available to physicians in the United States remains an open question.

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# Reference

- Poynard T, Naveneau S, Mory B, Chaput U. Meta-analysis of smooth muscle relaxants in the treatment of irritable bowel syndrome. Aliment Pharmacol Ther. 1994;8:499-510.
- Jailwala J, Imperiale T, Kroenke K. Pharmacologic treatment of the irritable bowel syndrome: a systematic review of randomized, controlled trials. Ann Intern Med. 2000;133:136-47.

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