

AGA-ACG Clinical Practice Guideline: Pharmacological Management of Chronic Idiopathic Constipation By Hima Veeramachaneni

- <u>Chronic idiopathic constipation (CIC</u>) = disorder of gut-brain interaction causing infrequent & incomplete defecation w/o mucosal or structural abnormalities
 Affects 8-12% of US population
 - Nonpharmacologic treatments = dietary changes (个 fluid intake & 个 dietary fiber), behavioral changes (i.e., exercise), squatty potty
- Pharmacologic treatments = fiber, osmotic laxatives, stimulant laxatives, secretagogues, serotonin type 4 agonist

**1st guideline to recommend MgO & senna as evidence-based treatments

Over-the-Counter (OTC) Pharmacologic Treatments for CIC <pre>vindicates strong recommendation</pre>											
Medication	Mechanism of Action	Initial dose	Maximum dose	Cost	Side Effects	Comments					
Fiber (1 st line for CIC)	Soluble = traps water to soften Insoluble = 个 bulk	14 g/ 1,000 kcal/ day	No benefit for total intake (dietary & supplement) > 25-30 g	\$	Flatulence	 -Need adequate hydration -Psyllium = only effective fiber per studies -No evidence if soluble vs insoluble is more effective 					
✓Polyethylene Glycol (PEG)	Osmotic laxative	17 g daily	No clear maximum	\$	Bloating, abdominal discomfort/cramping	-Can be used with fiber -Durable response over 6 months					
Magnesium Oxide (MgO)	Osmotic laxative	400-500 mg daily	No clear maximum (~1,000-1,500 mg/day)	\$		 -Caution with renal insufficiency (due to risk of hypermagnesemia) & pregnancy 					
Lactulose	Osmotic laxative	15 g daily	No clear maximum	\$	Bloating, flatulence	 -Significant diarrhea → ↑ Na & ↓K -Only osmotic agent studied in pregnancy -In studies, led to ↓ impactions & need for enemas 					
✓Bisacodyl & Picosulfate	Stimulant laxative	Bisacodyl 5mg daily	10 mg oral daily	\$	Abdominal discomfort/cramping	-Short-term use (<4 weeks) or rescue therapy -Prolonged/excess use → diarrhea, ↑/↓ electrolyte -Contraindicated in ileus/obstruction, severe dehydration, or acute inflammatory conditions					
Senna	Stimulant laxative	8.6-17.2 mg/daily	No clear maximum (~4 tabs, 2x daily)	\$	Abdominal discomfort/cramping	 Present in many laxative teas Long-term safety/efficacy unknown 					

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Reserved for those who do not respond to OTC therapy agents \rightarrow can be used as adjunct to OR replacement of OTC therapy All are contraindicated in known/suspected mechanical GI obstruction

Medication	Mechanism of Action	Initial dose	Maximum dose	Cost	Side Effects	Approved for IBS-C	Comments
Lubiprostone	Intestinal secretagogue acting on chloride channel type 2 → ↑ chloride secretion	24 μg BID	24 μg BID	\$\$	Dose- dependent nausea, diarrhea	Yes (8 μg BID dosing)	-Potential benefit for abdominal pain -Nausea risk is lower if taken with food & water -Adjust dosage for moderate/severe hepatic insufficiency (8 μg BID)
√ Linaclotide	Intestinal secretagogue acting on guanylate cyclase-C → activates CFTR → ↑ chloride secretion	72 – 145 μg daily	290 µg daily	\$\$\$	Diarrhea	Yes	-Potential benefit for abdominal pain -Most common reason for discontinuation in 1 st year of therapy= loss of efficacy & insurance barriers
√ Plecanatide	Intestinal secretagogue acting on guanylate cyclase-C → activates CFTR → ↑ chloride secretion	3 mg daily	3 mg daily	\$\$\$	Diarrhea	Yes	
✓Prucalopride	5-HT ₄ agonist	1-2 mg daily	2mg daily	\$\$\$	Diarrhea, abdominal pain, nausea, headache	No	 -Potential benefit for abdominal pain -Caution: unusual changes in mood/behavior, suicidal ideation -Contraindicated in intestinal perforation, IBD, & toxic megacolon/megarectum

√indicates strong recommendations

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