# **UW** Integrative Health

Department of Family Medicine and Community Health

## **Coming Off a Proton Pump Inhibitor**

#### Tapering

For patients who have made positive lifestyle changes and are less likely to need continued chronic acid suppression, it can still be difficult to come off PPIs. They often cause rebound hyperacidity, even if the underlying condition has resolved.<sup>1</sup> This occurs due to the lower stomach acidity increasing gastrin secretion, which causes the enterochromaffin cells to hypertrophy. When the PPI is suddenly discontinued, these larger cells have an increased capacity for acid secretion.<sup>2</sup> Figure 1 shows symptoms scores for dyspepsia in *asymptomatic* people given 40 milligrams of pantoprazole for 6 weeks versus controls. Despite being initially asymptomatic, they experienced rebound dyspepsia that lasted 10-14 days.<sup>1</sup>



Figure 1. Symptoms of reflux in people without GERD when taking PPIs versus placebo.<sup>1</sup> Blue, dashed = Took PPI; Red, solid = Placebo group. Reprinted by permission from Macmillan Publishers LTD: American Journal of Gastroenterology, copyright 2010.

When counseling about discontinuing a PPI, let patients know that they will likely have symptoms of reflux for about 2 weeks after they stop the medication. Fortunately, there are strategies to help calm reflux symptoms until rebound hyperacidity resolves.

#### **Bridge Therapy**

The following therapies will not only increase success for discontinuing a PPI but also are therapeutic for gastroesophageal reflux disease (GERD).

- 1. Focus on nutrition. Common foods that should be avoided in those with GERD include alcohol, caffeine (coffee), chocolate, cow's milk, animal fat, and orange juice.
- 2. Slowly taper off the PPI over 2-4 weeks (the higher the dose, the longer the taper).

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- 3. While the taper is being completed, use the following for bridge therapy to reduce the symptoms of rebound hyperacidity. Encourage regular aerobic exercise.
  - Encourage a relaxation technique such as deep breathing. This enhances vagal stimulation, encouraging digestion, and aids adequate peristalsis. or more information, see the "<u>Mind and Emotions</u>" and "<u>Mindful Awareness</u>" overviews.
  - Consider acupuncture 1-2 times per week.<sup>3</sup> Add one or more of the following dietary supplements: **Deglycyrrhizinated licorice** (DGL), 2-4 380 mg tablets before meals or sucralfate (Carafate) 1 gm before meals
    - **Slippery elm**, 1-2 tbsp of powdered root in water or 400-500 mg capsules or 5 mL of a tincture three to four times daily.
    - A combination botanical product, **Iberogast** 1 ml three times daily.<sup>4</sup>
- 4. If the patient is successful with stopping the PPI, slowly taper off the above (except for positive nutritional changes, exercise, and stress management). If symptoms return, start again with one of the above or an H2 blocker (e.g., Ranitidine, 150 mg twice daily or as needed). If symptoms are still difficult to control, consider adding the PPI back at the lowest effective dose.

Note: PPIs shut off all three acid pumps and H2 blockers are partial inhibitors of acid secretion. If long-term treatment is needed, H2 blockers allow better absorption of nutrients than PPIs and so potentially have fewer long-term adverse effects.

- 5. For those with reflux hypersensitivity (those with normal endoscopies, normal pH monitoring/physiologic reflux, no esophageal motor disorder), consider an SSRI as first-line and an SNRI or tricylic anti-depressant as second-line. Patients with this condition are also more likely to have co-morbid behavioral disorders.<sup>2</sup>
- 6. It would be most beneficial to avoid long-term acid suppression if possible since this is associated with malabsorption of vitamin B12<sup>5</sup> and iron,<sup>6</sup> increased risk of community-acquired pneumonia,<sup>7</sup> hip<sup>8, 9</sup> and spine<sup>10, 11</sup> fractures, *C. diff* diarrhea<sup>6, 12</sup>, and gastric cancer if a long-term PPI is used post H. pylori eradication therapy. <sup>13</sup> Early research has also suggested an association between chronic PPI use and dementia, which may be due to increases in amyloid plaque deposition.<sup>14, 15</sup> Finally, PPIs have been associated with higher rates of acute interstitial nephritis and end-stage renal disease, so it may be prudent to wean off of them in patients at risk for chronic kidney disease progression.<sup>15-17</sup> For more details, check out the "<u>Gastroesophageal Reflux Disease (GERD)</u>" tool.

### **Resource Links**

- <u>Mind and Emotions:</u> https://www.fammed.wisc.edu/integrative/resources/modules/mind-and-emotions/
- <u>Mindful Awareness</u>: https://www.fammed.wisc.edu/integrative/resources/modules/meditation/mindfulawareness/
- <u>Gastroesophageal Reflux Disease (GERD)</u>: https://www.fammed.wisc.edu/files/webfm-uploads/documents/outreach/im/tool-gerd

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This tool was adapted for the University of Wisconsin Integrative Health Program from the original, written by David Rakel, MD (2014) and updated by David Lessens, MD, MPH and Sagar Shah, MD (2020). Sections were adapted from a previous version of "Gastroesophageal Reflux Disease" written by David Kiefer, MD, David Rakel, MD, and Rian Podein, MD. Modified for UW Integrative Health in 2021.

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#### References

- Niklasson A, Lindstrom L, Simren M, Lindberg G, Bjornsson E. Dyspeptic symptom development after discontinuation of a proton pump inhibitor: a double-blind placebo-controlled trial. *Am J Gastroenterol.* 2010;doi:10.1038/ajg.2010.81
- 2. Kim J, Blackett JW, Jodorkovsky D. Strategies for effective discontinuation of proton pump inhibitors. *Curr Gastroenterol Rep.* May 16 2018;20(6):27. doi:10.1007/s11894-018-0632-y
- 3. Dickman R, Schiff E, Holland A, et al. Clinical trial: acupuncture vs. doubling the proton pump inhibitor dose in refractory heartburn. *Aliment Pharmacol Ther*. Nov 15 2007;26(10):1333-44. doi:10.1111/j.1365-2036.2007.03520.x
- 4. Melzer J, Rosch W, Reichling J, Brignoli R, Saller R. Meta-analysis: phytotherapy of functional dyspepsia with the herbal drug preparation STW 5 (Iberogast). *Aliment Pharmacol Ther*. Dec 2004;20(11-12):1279-87. doi:10.1111/j.1365-2036.2004.02275.x
- 5. Lam JR, Schneider JL, Zhao W, Corley DA. Proton pump inhibitor and histamine 2 receptor antagonist use and vitamin B12 deficiency. *Nat Med.* Dec 11 2013;310(22):2435-42. doi:10.1001/jama.2013.280490
- 6. Wilhelm SM, Rjater RG, Kale-Pradhan PB. Perils and pitfalls of long-term effects of proton pump inhibitors. *Expert Rev Clin Pharmacol.* Jul 2013;6(4):443-51. doi:10.1586/17512433.2013.811206
- Laheij RJ, Sturkenboom MC, Hassing RJ, Dieleman J, Stricker BH, Jansen JB. Risk of communityacquired pneumonia and use of gastric acid-suppressive drugs. *JAMA*. Oct 27 2004;292(16):1955-60. doi:10.1001/jama.292.16.1955
- Corley DA, Kubo A, Zhao W, Quesenberry C. Proton pump inhibitors and histamine-2 receptor antagonists are associated with hip fractures among at-risk patients. *Gastroenterology*. Jul 2010;139(1):93-101. doi:10.1053/j.gastro.2010.03.055
- Gray SL, LaCroix AZ, Larson J, et al. Proton pump inhibitor use, hip fracture, and change in bone mineral density in postmenopausal women: results from the Women's Health Initiative. Arch Intern Med. May 10 2010;170(9):765-71. doi:10.1001/archinternmed.2010.94
- 10. Kwok CS, Yeong JK, Loke YK. Meta-analysis: Risk of fractures with acid-suppressing medication. *Bone*. 2011;48(4):768-76. doi:10.1016/j.bone.2010.12.015
- 11. Insogna KL. The effect of proton pump-inhibiting drugs on mineral metabolism. *Am J Gastroenterol.* 2009;104:S2-S4.
- 12. Cunningham R, Dale B, Undy B, Gaunt N. Proton pump inhibitors as a risk factor for Clostridium difficile diarrhoea. *J Hosp Infect*. 2003;54(3):243-245. doi:S0195670103000884 [pii]
- Tan MC, Graham DY. Proton pump inhibitor therapy after Helicobacter pylori eradication may increase the risk of gastric cancer. *BMJ Evid Based Med.* Jun 2018;23(3):111-112. doi:10.1136/bmjebm-2018-110935
- Gomm W, von Holt K, Thomé F, et al. Association of proton pump inhibitors with risk of dementia: a pharmacoepidemiological claims data analysis. *JAMA Neurol.* Apr 2016;73(4):410-6. doi:10.1001/jamaneurol.2015.4791

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- 15. Klepser DG, Collier DS, Cochran GL. Proton pump inhibitors and acute kidney injury: a nested casecontrol study. *BMC Nephrol.* Jul 16 2013;14:150. doi:10.1186/1471-2369-14-150
- Antoniou T, Macdonald EM, Hollands S, et al. Proton pump inhibitors and the risk of acute kidney injury in older patients: a population-based cohort study. *CMAJ open*. Apr-Jun 2015;3(2):E166-71. doi:10.9778/cmajo.20140074
- 17. Lazarus B, Chen Y, Wilson FP, et al. Proton pump inhibitor use and the risk of chronic kidney disease. *JAMA Intern Med.* Feb 2016;176(2):238-46. doi:10.1001/jamainternmed.2015.7193

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