

# GUIDELINES & PROTOCOLS

## ADVISORY COMMITTEE

### Gastroesophageal Reflux Disease - Clinical Approach in Adults

Effective Date: January 30<sup>th</sup>, 2009

#### Scope

This guideline outlines the clinical approach to the diagnosis and treatment of gastroesophageal reflux disease (GERD) in adult patients. Treatment of *Helicobacter pylori* (*H. pylori*) infection is not part of the management of GERD (see Dyspepsia with or without *H. pylori* Infection guideline).

**Diagnostic Code:** 536 Dyspepsia; 535 or 537 Gastritis and Duodenitis

#### Prevention and Risk Factors

Obesity is a major risk factor.<sup>1,2</sup> Symptoms may be aggravated by spicy or fatty foods, caffeine, alcohol, citrus fruits, recumbency or bending forward.<sup>3,4</sup> GERD may also be provoked by certain medications such as calcium channel blockers and may be mimicked by other drugs such as bisphosphonates and non-steroidal anti-inflammatory drugs (NSAID).<sup>5</sup> GERD is frequently worse during pregnancy (see Management of GERD in pregnancy).

#### Diagnosis/Investigation

GERD is usually diagnosed by history. Symptoms typically include retrosternal burning and may also include sour or bilious regurgitation, belching, hypersalivation, and epigastric or chest pain.<sup>6</sup> Increasingly recognized are extraesophageal symptoms such as chronic cough, laryngeal irritation and wheezing, particularly when they occur at night.<sup>7,4</sup> Certain symptoms ('alarm features') require prompt endoscopy. These include dysphagia, weight loss, gastrointestinal blood loss (acute or chronic), persistent vomiting or failure to respond to an adequate trial of therapy.<sup>6,8</sup> Differential diagnoses to consider include cardiac and musculoskeletal disorders.

#### Management

##### Initial Management of GERD:

In the absence of alarm features the initial management should consist of diet and lifestyle modifications, antacids, alginates or histamine<sub>2</sub> receptor antagonists (H<sub>2</sub>RA) (see Appendix A).<sup>6,9</sup> Under these circumstances barium X-rays and endoscopy results are frequently normal and are generally not recommended.<sup>6</sup> Antacids and alginates may be effective in patients with intermittent or sporadic symptoms.

##### Management of severe symptoms or poor response:

In the absence of improvement with the above management strategy, H<sub>2</sub>RA or proton pump inhibitors (PPI) may be tried (see Appendix A). It may take 4-8 weeks to see a response. GERD is a chronic disease and patients may require prolonged or intermittent therapy.<sup>10</sup> H<sub>2</sub>RAs and PPIs are more effective in patients with chronic symptoms.

### Management of refractory symptoms:

Absence of response to the above regimens justifies specialist consultation and/or further investigation. Endoscopy is the investigation of choice.

### Management of GERD in pregnancy:

Traditional antacids and alginates are generally considered safe in pregnancy and lactation, and can be considered first-line in this setting. Studies on H<sub>2</sub>RA and PPI in pregnancy do not demonstrate an increased risk of malformations; these are appropriate second line agents.<sup>11,12</sup> In lactation, cimetidine is recognized as safe whereas other H<sub>2</sub>RA and PPI have not been adequately studied. For the latest information on drug safety in pregnancy and lactation, please refer to recognized database sources such as [www.motherisk.org](http://www.motherisk.org)

### **Rationale**

GERD is a common chronic recurrent problem. Most individuals with GERD experience only occasional heartburn, which is usually responsive to simple measures. GERD and hiatus hernia are not synonymous and do not imply each other's presence.

More severe reflux can cause esophageal mucosal injury (esophagitis) and its complications. Respiratory symptoms (chronic cough, hoarseness, bronchospasm, recurrent aspiration) may occur in the absence of typical heartburn. Patients with extraesophageal symptoms such as chronic cough may not respond well or quickly to standard antireflux therapy.<sup>4</sup>

Chronic longstanding GERD may be complicated by Barrett's esophagus (intestinal metaplasia in the lower esophagus) in up to 10% of individuals.<sup>13</sup> Barrett's esophagus predisposes to adenocarcinoma, with an incidence of 0.5-1% per year.<sup>14</sup> This risk for cancer is higher in caucasians, males, individuals aged >50 years, smokers, and people with more than 10 years of symptoms occurring more than 3 times per week.<sup>4</sup> Patients with the above risk factors may be offered endoscopy on one occasion to rule out Barrett's esophagus; if not present, it will generally not develop later.

Endoscopy is not necessary or universally effective in making a diagnosis of GERD, but is considered the investigation of choice to identify esophagitis, assess its severity and rule out complications including strictures and Barrett's esophagus. Barium studies are not adequate to assess the mucosa or diagnose reflux disease.<sup>4</sup>

Patients with complicated GERD (Barrett's esophagus, ulceration, bleeding, peptic stricture) may require long-term PPI therapy.<sup>10,15</sup> The efficacy of prokinetic agents (domperidone and metaclopramide) has not been established.

Anti-reflux surgery could be considered in patients who respond well to PPI therapy, but who are intolerant or reluctant to take medications. Outcomes are highly dependent on individual factors.<sup>10</sup>

### **References**

- <sup>1</sup> Hampel H, Abraham NS, El-Serag HB. Meta-analysis: obesity and the risk for gastroesophageal reflux disease and its complications. *Ann.Intern.Med.* 2005 Aug 2;143(3):199-211.
- <sup>2</sup> Jacobson BC, Somers SC, Fuchs CS, et al. Body-mass index and symptoms of gastroesophageal reflux in women. *N.Engl.J.Med.* 2006 Jun 1;354(22):2340-2348.
- <sup>3</sup> Kaltenbach T, Crockett S, Gerson LB. Are lifestyle measures effective in patients with gastroesophageal reflux disease? An evidence-based approach. *Arch.Intern.Med.* 2006 May 8;166(9):965-971
- <sup>4</sup> Richter JE. Gastroesophageal reflux disease. *Best Practice & Research CLinical Gastroenterology* 2007;21(4):609-631.
- <sup>5</sup> Leong R, Chan F. Drug-induced side effects affecting the gastrointestinal tract. 2006;5(4):585.

- 
- <sup>6</sup> Armstrong D, Marshall JK, Chiba N, et al. Canadian Consensus Conference on the management of gastroesophageal reflux disease in adults - update 2004. *Can.J.Gastroenterol.* 2005 Jan;19(1):15-35.
- <sup>7</sup> Chang AB, Lasserson TJ, Kiljander TO, et al. Systematic review and meta-analysis of randomised controlled trials of gastro-oesophageal reflux interventions for chronic cough associated with gastro-oesophageal reflux. *BMJ* 2006 Jan 7;332(7532):11-17.
- <sup>8</sup> Jones R. Gastro-oesophageal reflux disease: a re-appraisal. *Br.J.Gen.Pract.* 2006 Oct;56(531):739-740.
- <sup>9</sup> Tran T, Lowry AM, El-Serag HB. Meta-analysis: the efficacy of over-the-counter gastro-oesophageal reflux disease therapies. *Aliment.Pharmacol.Ther.* 2007 Jan 15;25(2):143-153.
- <sup>10</sup> Fennerty MB. Review article: alternative approaches to the long-term management of GERD. *Aliment.Pharmacol.Ther.* 2005 Dec;22 Suppl 3:39-44.
- <sup>11</sup> Magee L, Inocencion G, Kamboj L, et al. Safety of First Trimester Exposure to Histamine H2 Blockers. A Prospective Cohort Study. *Digestive Diseases and Sciences* 1996;41(6):1145.
- <sup>12</sup> Nikfar S, Mohammad D, Mula M, M., et al. Use of Proton Pump Inhibitors During Pregnancy and Rates of Major Malformations. A Meta-analysis. *Digestive Diseases and Sciences* 2002;47(7):1526.
- <sup>13</sup> Spechler S, Goyal R. The Columnar-Lined Esophagus, Intestinal Metaplasia, and Norman Barrett. *Gastroenterology* 1996;110:614.
- <sup>14</sup> Shaheen N, Ransohoff DF. Gastroesophageal reflux, barrett esophagus, and esophageal cancer: scientific review. *JAMA* 2002 Apr 17;287(15):1972-1981.
- <sup>15</sup> Wang KK, Sampliner RE, Practice Parameters Committee of the American College of Gastroenterology. Updated guidelines 2008 for the diagnosis, surveillance and therapy of Barrett's esophagus. *Am.J.Gastroenterol.* 2008 Mar;103(3):788-797.

## List of Abbreviations

GERD	Gastroesophageal reflux disease
NSAID	Non-steroidal anti-inflammatory drugs
H <sub>2</sub> RA	Histamine <sub>2</sub> receptor antagonist
PPI	Proton pump inhibitor

This guideline is based on scientific evidence current as of the Effective Date.

## Resources

[www.motherisk.org](http://www.motherisk.org)

## Contact Information

Guidelines and Protocols Advisory Committee  
PO Box 9642 STN PROV GOVT  
Victoria BC V8W 9P1

Telephone: (250) 952-1347  
Fax: (250) 952-1417

E-mail: [h1th.guidelines@gov.bc.ca](mailto:h1th.guidelines@gov.bc.ca)  
Web site: [www.BCGuidelines.ca](http://www.BCGuidelines.ca)

The principles of the Guidelines and Protocols Advisory Committee are to:

- encourage appropriate responses to common medical situations
- recommend actions that are sufficient and efficient, neither excessive nor deficient
- permit exceptions when justified by clinical circumstances

## Appendices

### Appendix A – Prescription Medication table for Gastroesophageal reflux disease

#### Appendix A – Prescription Medication table for Gastroesophageal reflux disease

Drug	Standard Rx	Approx cost per Standard Rx (Date)	PharmaCare Coverage
<b>H<sub>2</sub>-receptor antagonists (H<sub>2</sub>RA)</b>			
cimetidine (Tagamet®)	1200 mg per day in divided doses x 8-12 weeks	\$90 - \$249 G:\$22 - \$42	regular benefit, LCA
famotidine (Pepcid®)	20 mg bid*	\$132 [for 8 weeks] G: \$76 [for 8 weeks]	*
nizatidine (Axid®)	150 mg bid x 12 weeks	\$162 G: \$102	partial coverage, RDP, LCA
ranitidine (Zantac®)	150 mg bid x 8 weeks*	\$140-\$170 G: \$52	*
<b>Proton Pump inhibitors (PPI)</b>			
esomeprazole (Nexium®)	20-40 mg qd x 4-8 weeks	\$67 - \$134	limited coverage; For GERD, patient must have treatment failure or intolerance to cimetidine or other H <sub>2</sub> RA, plus failure of an 8-week trial or intolerance to rabeprazole.
lansoprazole (Prevacid®)	15-30 mg qd x 4-8 weeks	\$64 - \$128	
omeprazole (Losec®)	20 mg qd x 4 weeks	\$70 G: \$40	
pantoprazole (Pantaloc®)	40 mg qd x 4 weeks	\$63 G:\$44	limited coverage; For GERD, patient must have treatment failure or intolerance to cimetidine or other H <sub>2</sub> RA.
rabeprazole (Pariet®)	20 mg qd x 4 weeks	\$42 G: \$29	

**Nb:** Please review product monographs and regularly review current listings of Health Canada advisories, warnings and recalls at: [http://www.hc-sc.gc.ca/ahc-asc/media/advisories-avis/index\\_e.html](http://www.hc-sc.gc.ca/ahc-asc/media/advisories-avis/index_e.html)

See <http://www.health.gov.bc.ca/pharme/> for further information.

**G:** indicates that generics are available.

\* available at this dose without a prescription, but non-prescription medications are not reimbursed by PharmaCare or most private drug plans.

**Regular benefit drugs:** do not require Special Authority. Patients may receive full or partial coverage, since some of these drugs are included in the Low Cost Alternative (LCA) program or Reference Drug Program (RDP).

**LCA:** When multiple medications contain the same active ingredient (usually generic products), patients receive full coverage for the drug with the lowest average PharmaCare claimed price. The remaining products are partial benefits.

**RDP:** When a number of products contain different active ingredients but are in the same therapeutic class, patients receive full coverage for the drug that is medically effective and the most cost-effective. This drug is designated as the Reference Drug. The remaining products are partial benefits.

**Limited coverage drugs:** require Special Authority. These drugs are not normally regarded as first-line therapies or there are drugs for which a more cost-effective alternative exists.

**In all cases:** coverage is subject to drug price limits set by PharmaCare and to the patient's PharmaCare plan rules and deductibles.