



# Digestive Enzymes

©2010 Huntington College of Health Sciences

Literature Education Series On Dietary Supplements

By Gene Bruno, MS, MHS – Dean of Academics, Huntington College of Health Sciences

*Smart Supplementation™ is a free series of educational literature created by Huntington College of Health Sciences (HCHS) as a public service. Although copyrighted, it may be freely photocopied and distributed, but may not be altered in any way. Smart Supplementation™ is not intended as medical advice. For diagnosis and treatment of any medical condition, consult your physician.*

Do you have indigestion, heartburn, bloating or gassiness? If so, rather than blocking the digestive process with antacids, you may wish to try the natural approach which focuses on aiding digestion.

Digestive enzymes and hydrochloric acid supplements are often recommended by practitioners of natural medicine to help improve digestion and prevent reflux.<sup>1</sup>

## **Low digestive enzyme and hydrochloric acid secretion**

The rationale for this approach is that a more likely cause of indigestion is a lack of digestive enzyme secretion as well as a lack of hydrochloric acid secretion, or hypochlorhydria, rather than hyperacidity.<sup>2</sup> In fact, both human and animal research has demonstrated that digestive enzymes produced by the pancreas reduce with age.<sup>3,4</sup> In addition, suboptimal pancreatic function may occur for other reasons, including pancreatic and non-pancreatic disorders—both of which may cause an impaired production of pancreatic digestive enzymes, resulting in poor digestion. At least in part, this is there is also an age-related decline in bile acid synthesis by the liver<sup>5</sup>—bile acids help digest fats. Furthermore, insufficient amounts of digestive enzymes can cause or exacerbate abnormal digestive conditions, such as maldigestion, food allergies or sensitivities, intestinal fermentation, putrefaction and peroxidation, and the phenomenon known as intestinal hyperpermeability, or “leaky gut.”<sup>6</sup>

Likewise, several studies have shown that the ability to secrete hydrochloric acid decreases with age<sup>7,8,9</sup>,

with low stomach acidity in more than half of the subjects older than 60 years in some cases. In addition some research suggests that people with a wide variety of chronic disorders do not produce adequate amounts of stomach acid. These disorders include allergies<sup>10,11,12</sup>, asthma<sup>13,14</sup>, gallstones<sup>15,16</sup>, rosacea<sup>17</sup>, dermatitis herpetiformis<sup>18</sup>, rheumatoid arthritis<sup>19</sup>, and vitiligo.<sup>20</sup>

Inadequate secretion of digestive enzymes and hydrochloric acid can be addressed through the use of a multi-digestive enzyme (MDE) supplement. Such supplements may contain pancreatin or pancreatic lipase, pancreas extracts from pork (porcine) or beef (bovine) sources that contain lipase, protease, and amylase.<sup>21</sup> They may also contain microbial/fungal sources of enzymes, as well as betaine hydrochloride as a source of hydrochloric acid, and ox bile as a source of bile.

## **Digestive enzymes**

Many, though not all, of digestive enzymes are produced by the pancreas, and each has specific functions. *Lipase* is a digestive enzyme that is widely distributed in the plant world, in milk, milk products, bacteria, molds, and animal tissues.<sup>22</sup> Lipase enzymes aid in fat digestion by hydrolyzing fat in the small intestine. *Amylase* is an enzyme that breaks starch down into sugar. *Protease* is any enzyme that conducts proteolysis, that is, begins protein catabolism by hydrolysis of the peptide bonds that link amino acids together in the polypeptide chain, which form a molecule of protein.

## **Using Pancreatin or Pancrealipase**

Pancreatin or pancreatic lipase (a more concentrated form of pancreatin) has been successful in improving digestion function. In one study, previously housebound patients with suboptimal pancreatic function who used pancreatin were able to return to a near-normal social and work life-style.<sup>23</sup> In another study, patients with impaired digestion due to severe, suboptimal pancreatic function also experienced impaired secretion of cholecystokinin (CCK)<sup>24</sup>—an intestinal hormone that stimulates bile secretion and

consequent fat digestion. Supplementation with pancreatin caused these patients to experience a significant increase in their CCK levels.<sup>25</sup>

A number of studies have shown that patients with impaired digestion due to suboptimal pancreatic function experienced improved digestion after supplementation with pancreatin.<sup>26 27 28</sup> Research has also shown that individuals with chronic suboptimal pancreas function also experienced digestive improvement with pancreatin.<sup>29 30</sup> In addition, several studies have shown that inadequate digestive enzyme production can be caused by various non-pancreatic disorders, and responds well to digestive enzyme supplementation.<sup>31</sup>

Since pancreatin is a good source of lipase, it is no surprise that a pancreatin supplement reduced fat in the stools and improved nutritional status in patients with suboptimal pancreatic function.<sup>32 33 34</sup> Likewise, in a double-blind study lipase from pancreatin was shown to significantly reduce gas, bloating, and fullness after a high-fat meal.<sup>35</sup> Participants in this study took one capsule immediately before the meal and two capsules immediately after the meal.

Because pancreatin is rapidly emptied from the stomach during digestion, people taking digestive enzyme product may obtain better results by spreading out supplementation throughout the meal.<sup>36</sup>

### **Microbial enzymes**

Inadequate digestive enzyme function can be addressed through the use of microbial and fungal enzymes with activities similar to pancreatin. Microbial enzymes have been the subject of various studies evaluating their effects on lactose intolerance, impaired pancreatic enzyme production, excess fat in the feces, celiac disorder and a variety of other digestive issues, with positive results.<sup>37 38 39 40 41 42</sup> In addition, a number of studies have shown that microbial enzymes have anti-inflammatory activity as well as fibrinolytic properties, as demonstrated by their ability to hydrolyze fibrin and fibrinogen<sup>43 44</sup> (found in scar tissue).

One of the most functionally valuable attributes related to microbial enzymes is that they appear to possess unusually high stability and activity throughout a wide range of pH conditions (from a pH of 2-10).<sup>45</sup> This enables them to be more consistently active and functional for a longer distance as they are transported through the digestive tract.

### **Betaine hydrochloride**

To address hypochlorhydria, the use of betaine hydrochloride as a supplemental source of hydrochloric acid is indicated. Based upon clinical experience<sup>46</sup>, this supplemental source of hydrochloric acid often relieves the symptoms of

heartburn and improve digestion in people who have hypochlorhydria.

Furthermore, the importance of maintaining healthy levels of hydrochloric acid is underscored by other key roles that this digestive aid plays in the body. For example, hydrochloric-acid secretion from the stomach, pancreatic enzymes, and bile all inhibit the overgrowth of *Candida* and prevent its penetration into the absorptive surfaces of the small intestine.<sup>47 48</sup> Likewise, many minerals and vitamins require adequate concentrations of stomach acid to be optimally absorbed from food or supplements.<sup>50 51 52 53 54 55</sup>

In addition, clinical improvements have been noted in the case of individuals with disorders associated with hypochlorhydria. In a preliminary trial, supplemental hydrochloric acid, along with B-complex improved some cases of rosacea<sup>56</sup> and hives<sup>57</sup> in people with low stomach-acid production. Supplementation with hydrochloric acid in combination with avoidance of known food allergens led to clinical improvement in children with asthma.<sup>58</sup> Also, supplementation with hydrochloric acid resulted in gradual repigmentation of the skin in some patients with vitiligo after one year or more.<sup>59</sup>

### **Pepsin**

Pepsin is an enzyme that degrades food proteins into peptides. In the stomach, chief cells release pepsinogen, a pro-form zymogen. Hydrochloric acid activates pepsinogen to convert to pepsin. Pepsin is stored as pepsinogen so it will only be released when needed, and does not digest the body's own proteins in the stomach's lining. Pepsin plays a complementary role to hydrochloric acid in digestion by facilitating the breakdown of proteins<sup>60</sup>, and rendering calcium and iron more absorbable.<sup>61</sup>

### **What are Bile Acids?**

Bile acids are the main active components of bile. Bile refers to the fluid produced by the liver and secreted into the small intestine from the gall bladder via the bile duct. Bile salt is a synonymous term for bile acid, and the two are often used interchangeably.<sup>62</sup> In a healthy person, bile acid synthesis is about 400 mg/day. However, bile acid secretion, is about 12,000 mg (12 grams) per day. The difference comes from bile acids which have been reabsorbed and reused within the small intestine.<sup>63</sup> As a consequence of this active reabsorption, a bile acid "pool" is formed that circulates multiple times daily. This gives an idea of how important bile acids are in the normal metabolism of the gastrointestinal tract.

In the small intestine, bile acids solubilize fats to aid their absorption. Deficiencies of bile acids cause fat malabsorption and fatty stools (steatorrhea) as indicated by diarrhea. In addition, bile acid

deficiency jeopardizes a person's nutritional status by reducing the absorption of fat and fat-soluble nutrients.<sup>64</sup> Bile acids have been shown in clinical trials to be effective in improving fat absorption and the nutritional status of individuals.<sup>65 66</sup>

### Regularity, Fatty Stool, Parasites

Bile acids have other roles besides the digestion of fats and fat-soluble nutrients. For instance, it softens the stool and encourages peristalsis, thus improving the movement of the digested food mass through the gastrointestinal tract. Diets that are high in soluble and semi-soluble fibers work with the liver to trap the bile in a form which will not be too quickly reabsorbed by the system. Furthermore, bile is one of the body's chief weapons against the colonization of parasites in the intestines.<sup>67</sup>

### References

<sup>1</sup> Golan R. Optimal Wellness. New York: Ballantine Books; 1995:373-4.  
<sup>2</sup> Pizzorno JE, Murray MT. Textbook of Natural Medicine, 3<sup>rd</sup> ed. Edinburgh: Churchill Livingstone; 2006.  
<sup>3</sup> Laugier R, et al. Changes in pancreatic exocrine secretion with age: pancreatic exocrine secretion does decrease in the elderly. *Digestion* 1991;50(3-4):202-11.  
<sup>4</sup> Wang CS, Floyd RA, Kloer HU. Effect of aging on pancreatic lipolytic enzymes. *Pancreas* 1986;1(5):438-42.  
<sup>5</sup> Einarsson K, Nilsell K, Leijd B, Angelin B. Influence of age on secretion of cholesterol and synthesis of bile acids by the liver. *NEJM* 1985; 313(5):277-282.  
<sup>6</sup> Rachman B. Unique features and application of non-animal derived enzymes. *Clinical Nutrition Insights* 1997; 5(10):1-4.  
<sup>7</sup> Rafsky HA, Weingarten M. A study of the gastric secretory response in the aged. *Gastroenterology* 1947;May:348-352.  
<sup>8</sup> Davies D, James TG. An investigation into the gastric secretion of a hundred normal persons over the age of sixty. *Br J Med* 1930;i:1-14.  
<sup>9</sup> Baron JH. Studies of basal and peak acid output with an augmented histamine test. *Gut* 1963;4:136-144.  
<sup>10</sup> Kokkonen J, Simila S, Herva R. Impaired gastric function in children with cow's milk intolerance. *Eur J Pediatr* 1979;132:1-6.  
<sup>11</sup> Kokkonen J, Simila S, Herva R. Gastrointestinal findings in atopic children. *Eur J Pediatr* 1980;134:249-54.  
<sup>12</sup> Gonzalez H, Ahmed T. Suppression of gastric H2-receptor mediated function in patients with bronchial asthma and ragweed allergy. *Chest* 1986;89:491-6.  
<sup>13</sup> Gillespie M. Hypochlorhydria in asthma with specific reference to the age incidence. *Q J Med* 1935;4:397-405.  
<sup>14</sup> Bray GW. The hypochlorhydria of asthma in childhood. *Q J Med* 1931;24:181-97.  
<sup>15</sup> Fravel RC. The occurrence of hypochlorhydria in gall-bladder disease. *Am J Med Sci* 1920;159:512-7.  
<sup>16</sup> Capper WM, Butler TJ, Kilby JO, Gibson MJ. Gallstones, gastric secretion and flatulent dyspepsia. *Lancet* 1967;i:413-5.  
<sup>17</sup> Johnson L, Eckardt R. Rosacea keratitis and conditions with vascularization of the cornea treated with riboflavin. *Arch Ophthalmol* 1940;23:899-907.  
<sup>18</sup> Yancy KB, Lawley TJ. "Immunologically Mediated Skin Diseases." Harrison's Online. 1999. <http://www.harrisonsonline.com/hill-bin/Chapters.cgi> (Jan 10, 2000).  
<sup>19</sup> Hartung EF, Steinbroker O. Gastric acidity in chronic arthritis. *Ann Intern Med* 1935;9:252.  
<sup>20</sup> Francis HW. Achlorhydria as an etiological factor in vitiligo, with report of four cases. *Nebraska State Med J* 1931;16(1):25-6.  
<sup>21</sup> McKevey GK, ed. AHFS Drug Information. Bethesda, MD: American Society of Health-System Pharmacists, 1998.

<sup>22</sup> Tursi JM, Phair PG, Barnes GL. Plant sources of acid stable lipases: potential therapy for cystic fibrosis. *J Paediatr Child Health* 1994;30:539-43.  
<sup>23</sup> Valerio D, et al. Clinical effectiveness of a pancreatic enzyme supplement. *JPEN J Parenter Enteral Nutr* 1981; 5(2):110-4.  
<sup>24</sup> Jansen JB, et al. Effect of pancreatic enzyme supplementation on postprandial plasma cholecystokinin secretion in patients with pancreatic insufficiency. *Regul Pept* 1989; 25(3):333-42  
<sup>25</sup> Jansen JB, et al. Effect of pancreatic enzyme supplementation on postprandial plasma cholecystokinin secretion in patients with pancreatic insufficiency. *Regul Pept* 1989; 25(3):333-42  
<sup>26</sup> Hillel PG, et al. The use of dual-isotope imaging to compare the gastrointestinal transit of food and pancreatic enzyme pellets in cystic fibrosis patients. *Nucl Med Commun* 1998; 19(8):761-9.  
<sup>27</sup> Benabdeslam H, et al. Biochemical assessment of the nutritional status of cystic fibrosis patients treated with pancreatic enzyme extracts. *Am J Clin Nutr* 1998; 67(5):912-8.  
<sup>28</sup> Ansaldi-Balocco N, Santini B, Sarchi C. Efficacy of pancreatic enzyme supplementation in children with cystic fibrosis: comparison of two preparations by random crossover study and a retrospective study of the same patients at two different ages. *J Pediatr Gastroenterol Nutr* 1988; 7 Suppl 1:S40-5.  
<sup>29</sup> Van Hoozen CM, et al. Efficacy of enzyme supplementation after surgery for chronic pancreatitis. *Pancreas* 1997; 14(2):174-80.  
<sup>30</sup> Delhay M et al. Comparative evaluation of a high lipase pancreatic enzyme preparation and a standard pancreatic supplement for treating exocrine pancreatic insufficiency in chronic pancreatitis. *Eur J Gastroenterol Hepatol* 1996; 8(7):699-703.  
<sup>31</sup> Gullo L. Indication for pancreatic enzyme treatment in non-pancreatic digestive diseases. *Digestion* 1993; 54 Suppl 2:43-7.  
<sup>32</sup> Thomson M, Clague A, Cleghorn GJ, Shepherd RW. Comparative in vitro and in vivo studies of enteric-coated pancrelipase preparations for pancreatic insufficiency. *J Pediatr Gastroenterol Nutr* 1993;17:407-13.  
<sup>33</sup> Owen G, Peters TJ, Dawson S, Goodchild MC. Pancreatic enzyme supplement dosage in cystic fibrosis. *Lancet* 1991;338:1153.  
<sup>34</sup> Stern RC, Eisenberg JD, Wagener JS, et al. A comparison of the efficacy and tolerance of pancrelipase and placebo in the treatment of steatorrhea in cystic fibrosis patients with clinical exocrine pancreatic insufficiency. *Am J Gastroenterol* 2000;95:1932-8.  
<sup>35</sup> Suarez F, Levitt MD, Adshear J, Barkin JS. Pancreatic supplements reduce symptomatic response of healthy subjects to a high fat meal. *Dig Dis Sci* 1999;44:1317-21.  
<sup>36</sup> Taylor CJ, Hillel PG, Ghosal S, et al. Gastric emptying and intestinal transit of pancreatic enzyme supplements in cystic fibrosis. *Arch Dis Child* 1999;80:149-52.  
<sup>37</sup> Alternative Medicine, the Definitive Guide. Future Medicine Publishing: Puyallup, WA. 1993;215-22.  
<sup>38</sup> Griffin SM, et al. Acid resistant lipase as replacement therapy in chronic pancreatic exocrine insufficiency: a study in dogs. *Gut* 1989;30:1012-15.  
<sup>39</sup> Schneider MU, et al. Pancreatic enzyme replacement therapy: comparative effects of conventional and enteric-coated microspheric pancreatin and acid-stable fungal enzyme preparations on steatorrhea in chronic pancreatitis. *Hepato-gastroenterol* 1985;32:97-102.  
<sup>40</sup> Rosado JL, et al. Enzyme replacement therapy for primary adult lactase deficiency. *Gastroenterol* 1984;87:1072-82.  
<sup>41</sup> Barillas C, Solomons NW. Effective reduction of lactose maldigestion in preschool by direct addition of beta-galactosidases to milk at mealtime. *Pediatrics* 1987;79(5):766-72.  
<sup>42</sup> Phelan JJ, et al. Celiac disease: the abolition of gliadin toxicity by enzymes from *Aspergillus niger*. *Clin Sci & Mol Med* 1977;53:35-43.  
<sup>43</sup> Selezneva AA, Bol'shakova MD. Proteolytic complex from *Aspergillus terricola*. *Prikl Biokhim Mikrobiol* 1986;22(1):3-11.  
<sup>44</sup> Selezneva AA, Babenko GA, Bol'shakova MD, Rozhanskaia TI, Margolina NA. Preparative isolation of terrilytin components and study of their enzymatic properties. *Prikl Biokhim Mikrobiol* 1976;12(3):416-20.

- <sup>45</sup> Griffin SM, et al. Acid resistant lipase as replacement therapy in chronic pancreatic exocrine insufficiency: a study in dogs. *Gut* 1989;30:1012-15.
- <sup>46</sup> Wright JV. Dr. Wright's Guide to Healing with Nutrition. New Canaan, CT: Keats Publishing, 1990, 155.
- <sup>47</sup> Boero M, Pera A, Andriulli A, et al. Candida overgrowth in gastric juice of peptic ulcer subjects on short- and long-term treatment with H2-receptor antagonists. *Digestion* 1983;28:158-63.
- <sup>48</sup> Rubinstein E. Antibacterial activity of the pancreatic fluid. *Gastroenterology* 1985;88:927-32 [review].
- <sup>49</sup> Sarker SA, Gyr R. Non-immunological defense mechanisms of the gut. *Gut* 1990;33:1331-7 [review].
- <sup>50</sup> Murray MJ, Stein N. A gastric factor promoting iron absorption. *Lancet* 1968;1:614.
- <sup>51</sup> Sturniolo GC, Montino MC, Rossetto L, et al. Inhibition of gastric acid secretion reduces zinc absorption in man. *J Am Coll Nutr* 1991;10:372-5.
- <sup>52</sup> Allison JR. The relation of hydrochloric acid and vitamin B complex deficiency in certain skin conditions. *South Med J* 1945;38:235-41.
- <sup>53</sup> Russell RM, Krasinski SD, Samloff IM. Correction of impaired folic acid (Pte Glu) absorption by orally administered HCl in subjects with gastric atrophy. *Am J Clin Nutr* 1984;39:656.
- <sup>54</sup> Schade SG, Cohen RJ, Conrad ME. Effect of hydrochloric acid on iron absorption. *N Engl J Med* 1968;279:672-4.
- <sup>55</sup> Bezwoda W, Charlton R, Bothwell T, et al. The importance of gastric hydrochloric acid in the absorption of nonheme food iron. *J Lab Clin Med* 1978;92:108-16.
- <sup>56</sup> Allison JR. The relation of hydrochloric acid and vitamin B complex deficiency in certain skin diseases. *South Med J* 1945;38:235-41.
- <sup>57</sup> Rawls WB, Ancona VC. Chronic urticaria associated with hypochlorhydria or achlorhydria. *Rev Gastroenterol* 1951;18:267-71.
- <sup>58</sup> Bray GW. The hypochlorhydria of asthma in childhood. *Q J Med* 1931;24:181-97.
- <sup>59</sup> Francis HW. Achlorhydria as an etiological factor in vitiligo, with report of four cases. *Nebraska State Med J* 1931;16(1):25-6.
- <sup>60</sup> Cox M, Nelson DR. *Lehninger Principles of Biochemistry*, 5<sup>th</sup> ed.. San Francisco: W. H. Freeman; 2008:96
- <sup>61</sup> Tso P, Crissinger K. Digestion and absorption of lipids. In Stipanuk MH, ed. *Biochemical and physiological aspects of human nutrition*. Philadelphia: WB Saunders, 2000:125-140.
- <sup>62</sup> Thibodeau GA, Patton KT. *Anatomy & Physiology*, 5<sup>th</sup> ed. St. Louis: Mosby; 2003.
- <sup>63</sup> Bowen R. *Pathophysiology of the Digestive System* (a hypertext). University of Colorado. Updated November 23, 2001. Retrieved May 20, 2009 from <http://www.vivo.colostate.edu/hbooks/pathphys/digestion/liver/bile.html>.
- <sup>64</sup> Heuther SE, McCance KL. *Understanding Pathophysiology*, 3<sup>rd</sup> ed. St. Louis: Mosby; 2004.
- <sup>65</sup> Kapral C, Wewalka F, Praxmarer V, Lenz K, Hofmann AF. Conjugated bile acid replacement therapy in short bowel syndrome patients with a residual colon. *Z Gastroenterol* 2004;42(7):583-9.
- <sup>66</sup> Heydorn S, Jeppesen PB, Mortensen PB. Bile acid replacement therapy with cholylsarcosine for short-bowel syndrome. *Scand J Gastroenterol* 1999;34(8):818-23.
- <sup>67</sup> Heuther SE, McCance KL. *Understanding Pathophysiology*, 3<sup>rd</sup> ed. St. Louis: Mosby; 2004.



For more than two decades, Huntington College of Health Sciences (HCHS) has offered more than a conventional undergraduate or graduate education. Our accredited\*, distance learning degrees and diploma programs also include the breadth of responsible complementary and alternative medicine viewpoints, providing our students with a well-rounded and comprehensive approach to nutrition and the health sciences:

- Master of Science in Nutrition
- Bachelor of Health Science in Nutrition
- Associate of Science in Applied Nutrition
- Diploma in Comprehensive Nutrition
- Diploma in Dietary Supplement Science
- Diploma in Sports Nutrition
- Diploma in Women's Nutrition
- Diploma in Natural Sciences
- Diploma in Small Business Management

1204D Kenesaw  
 Knoxville, TN 37919  
 865-524-8079 • 800-290-4226  
 E-Mail: [studentservices@hchs.edu](mailto:studentservices@hchs.edu)  
[www.hchs.edu.com](http://www.hchs.edu.com)

*\*Accredited member Distance Education & Training Council.*