

**Wyeth Animal Health**

400 Michener Road  
Guelph, Ontario  
N1K 1E4

**Wyeth**

April 7, 2005

Dear Doctor:

As you may be aware, the Veterinary Drugs Directorate (VDD) has issued a letter to veterinarians and animal health organizations regarding ProHeart<sup>®</sup> 6 (*a copy has been attached for your convenience*). Wyeth Animal Health feels it is important to address this letter directly and reiterate its complete confidence in the safety and efficacy of this unique heartworm preventative. Since its launch in 2002, Wyeth Animal Health has stood firmly behind ProHeart 6 and believes the product has proven to be of significant benefit to both veterinarians and pet owners in the prevention of potentially deadly heartworm disease.

In its letter, the VDD requested Wyeth Animal Health provide evidence to establish the safety of ProHeart 6 by May 2, 2005. Wyeth is responding promptly to this request and is confident the extensive and rigorously reviewed product data, as well as independent reports by scientific experts in veterinary medicine, will demonstrate the solid safety and efficacy profile of ProHeart 6, and will clearly address the VDD's questions about this product.

Specifically, Wyeth Animal Health data will show:

- The number of unfiltered adverse event reports, deemed possibly or probably related by the VDD, is 147, which equates to five reports per 10,000 doses sold in Canada.
- The number of allergic reports deemed probably related by the VDD (no concurrent vaccines) is two per 10,000 doses sold.
- The VDD's concern about hematologic, seizure and liver issues is not supported by Canadian data.
- There are only two possible hematologic and two seizure reports in the Canadian database, which represents a rate of 0.07 reports per 10,000 doses sold, which is deemed extremely rare by World Health Organization guidelines and represents a rate of disease below that normally seen in the canine population.
- The VDD also notes injury to the liver as a concern with ProHeart 6. Yet, there are no reports in the VDD's database categorized as part of the hepatobiliary system.



Part of Wyeth's submission to the VDD will also include a retrospective epidemiological study conducted by Lawrence Glickman, VMD, DrPh, FACE, Professor of Epidemiology and Public Health, and Head of the Clinical Epidemiology Section, School of Veterinary Medicine, Purdue University. He recently analyzed data from nearly seven million canine visits at more than 400 U.S. hospitals in the Banfield, The Pet Hospital<sup>®</sup> practice. Dr. Glickman's study concluded the safety profile of ProHeart 6 was similar to two commonly used monthly oral heartworm preventatives, and provided no support for the voluntary recall of the product from the U.S. veterinary market.

While Fort Dodge Animal Health, our U.S. counterpart, voluntarily complied with the U.S. Food and Drug Administration Center for Veterinary Medicine's (CVM) request to recall ProHeart 6 from the U.S. veterinary market, the company strongly disagreed with the Agency's interpretation of the data and the conclusions that were drawn. Fort Dodge is working closely with the CVM to address its remaining concerns and is optimistic this matter will be resolved successfully in the near future.

Since its launch in North America, more than 18 million doses of ProHeart have been safely and effectively administered to a wide variety of healthy dogs - protecting not only the individual dog to which it is administered, but also providing the veterinary profession with an important medicine to reduce the incidence of heartworm disease in the canine population. Our first and foremost concern is providing safe and effective products to ensure the health and well being of all pets.

Wyeth Animal Health appreciates the opportunity to address this matter with the VDD and looks forward to this product continuing to protect the canine population against heartworm disease. If you have questions about this matter, please call our Professional Services department at 1-800-267-1777. Thank you for your attention to this important information and for your support of ProHeart 6.

Sincerely,

A handwritten signature in black ink, appearing to read 'Lois Valli', is positioned above the typed name.

Lois Valli, B.Sc., M.Sc., D.V.M.  
Manager of Professional Services  
Wyeth Animal Health