

# Pimobendan & Degenerative Valve Disease: An EPIC Update

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## In the Literature

Boswood A, Häggström J, Gordon SG, et al. Effect of pimobendan in dogs with preclinical myxomatous mitral valve disease and cardiomegaly: the EPIC study—a randomized clinical trial. *J Vet Intern Med.* 2016;30(6):1765-1779.

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## FROM THE PAGE ...

Myxomatous mitral valve disease (MMVD), a common heart disease in dogs, is characterized by a long preclinical phase, and no therapies have been definitively proven to prolong this time before onset of congestive heart failure (CHF).

Dogs ( $n = 360$ ) were enrolled in this study (Evaluation of Pimobendan In dogs with Cardiomegaly caused by preclinical mitral valve disease [EPIC]) to compare the effectiveness of pimobendan vs placebo in delaying onset of CHF signs, cardiac-related death, or euthanasia. Dogs (9.1-33.1 lb [4.1-15 kg] body weight) were admitted if they were diagnosed with MMVD based on the presence of a grade 3/6 or greater left apical systolic murmur, echocardiographic evidence of degenerative changes to the mitral valve apparatus, and presence of mitral regurgitation. They also had to have sufficient cardiomegaly meeting specific echocardiographic and radiographic parameters for left atrial and left ventricular dilatation (**Figures 1**, next page, and **2**, page 57).

Dogs were administered pimobendan or placebo at 0.4-0.6 mg/kg PO, divided and given at 12-hour intervals. Each patient was determined to have reached the end of the study (ie, primary endpoint) if diagnosed with CHF via thoracic radiography or if they died or were euthanized due to cardiac disease.

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## Amoxicillin Trihydrate and Clavulanate Potassium Tablets

ANADA 200-592, Approved by FDA

**CAUTION:** Federal (USA) law restricts this drug to use by or on the order of a licensed veterinarian.

**INDICATIONS:** Amoxicillin Trihydrate and Clavulanate Potassium Tablets are indicated in the treatment of:

Dogs: Skin and soft tissue infections such as wounds, abscesses, cellulitis, superficial/juvenile and deep pyoderma due to susceptible strains of the following organisms:  $\beta$ -lactamase-producing *Staphylococcus aureus*, non- $\beta$ -lactamase-producing *Staphylococcus aureus*, *Staphylococcus* spp., *Streptococcus* spp., and *E. coli*.

Periodontal infections due to susceptible strains of both aerobic and anaerobic bacteria. Amoxicillin Trihydrate and Clavulanate Potassium Tablets have been shown to be clinically effective for treating cases of canine periodontal disease.

Cats: Skin and soft tissue infections such as wounds, abscesses, and cellulitis/dermatitis due to susceptible strains of the following organisms:  $\beta$ -lactamase-producing *Staphylococcus aureus*, non- $\beta$ -lactamase-producing *Staphylococcus aureus*, *Staphylococcus* spp., *Streptococcus* spp., *E. coli*, and *Pasteurella* spp.

Urinary tract infections (cystitis) due to susceptible strains of *E. coli*.

Therapy may be initiated with Amoxicillin Trihydrate and Clavulanate Potassium Tablets prior to obtaining results from bacteriological and susceptibility studies.

A culture should be obtained prior to treatment to determine susceptibility of the organisms to Amoxicillin Trihydrate and Clavulanate Potassium Tablets. Following determination of susceptibility results and clinical response to medication, therapy may be reevaluated.

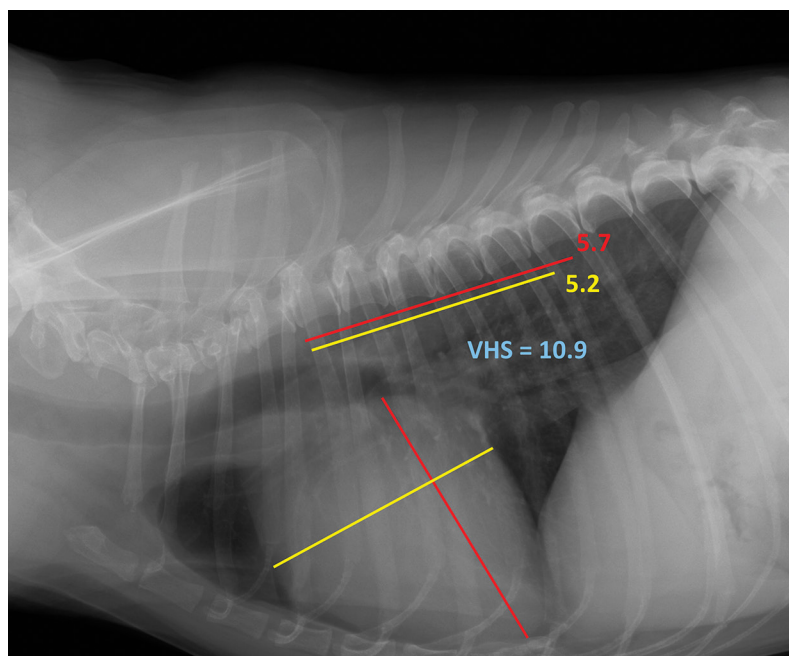
**CONTRAINDICATIONS:** The use of this product is contraindicated in animals with a history of an allergic reaction to any of the penicillins or cephalosporins.

**WARNINGS:** Safety of use in pregnant or breeding animals has not been determined.

**ADVERSE REACTIONS:** Amoxicillin Trihydrate and Clavulanate Potassium Tablets contain a semisynthetic penicillin (amoxicillin) and have the potential for producing allergic reactions. If an allergic reaction occurs, administer epinephrine and/or steroids.



Manufactured for:  
Putney, Inc., a wholly owned subsidiary of Dechra Pharmaceuticals, PLC.  
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Made in Austria.

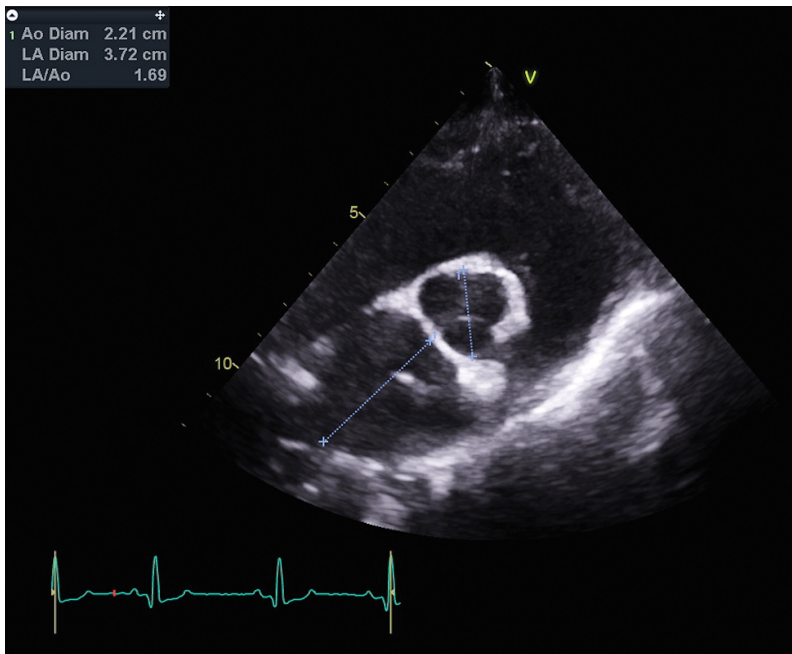


▲ **FIGURE 1** Measurement of VHS on a patient diagnosed with degenerative valve disease (via echocardiogram) that would meet the criteria for radiographic evidence of cardiomegaly according to the EPIC study criteria (ie, VHS >10.5)

An interim analysis demonstrated a clear treatment benefit for patients in the pimobendan group, so the study was terminated early. The pimobendan group had a median time to primary endpoint of 1228 days as compared with 766 days for the placebo group. There also was no difference in adverse events, indicating pimobendan was well tolerated with minimal side effects (the most common being vomiting and diarrhea).

This study demonstrated the benefit of pimobendan in significantly prolonging the time to onset of CHF or cardiac-related death in dogs with cardiomegaly due to MMVD.

**Pimobendan was well tolerated with minimal side effects.**



▲ **FIGURE 2** Measurement of the left atrial:aortic root ratio on a patient diagnosed with degenerative valve disease that would meet the criteria for left atrial enlargement according to the EPIC study criteria (ie, LA/Ao >1.6)

## ... TO YOUR PATIENTS

Key pearls to put into practice:

- 1** In patients with suspected MMVD based on the presence of a left apical systolic murmur in a middle-aged to older dog, thoracic radiographs are a beneficial first line to assess for cardiomegaly using vertebral heart size (VHS) measurement.
- 2** Echocardiography is essential to confirm diagnosis of MMVD and give detailed assessment of specific left atrial and left ventricular chamber enlargement criteria.
- 3** Chronic oral therapy with pimobendan should be initiated at 0.4-0.6 mg/kg (divided into q12h dosing) in patients without clinical signs that are diagnosed with cardiomegaly due to MMVD.

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