

IMPORTANT NEWS FOR US “BOXER BREEDERS”

Vet cardiologist discovers gene for heart disease

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Contact: Charlie Powell, WSU College of Veterinary Medicine, 509-335-7073, cpowell@vetmed.wsu.edu



PULLMAN — WSU veterinary cardiologist **Kathryn M. Meurs** discovered a mutant gene in the Boxer breed that causes a type of heart disease that can be fatal in animals and humans.

Well known in the Boxer breed community, the disease is called Boxer cardiomyopathy. The more formal term is arrhythmogenic right ventricular cardiomyopathy or ARVC.

This is same type of heart disease that caused the sudden death of 1950s college and pro football great Joe Campanella at age 36, as he played handball with the new head coach of the Baltimore Colts, Don Shula.

In Boxers, the disease can be fatal and frequently occurs when the animals exercise or become excited. Occasionally, they perish from the disease while at rest, too.

“Dr. Meurs’ discovery of both the gene and its location is a tremendous achievement in the cardiology of humans and animals,” said **Bryan Slinker**, dean of WSU’s College of Veterinary Medicine, and a recognized cardiac disease researcher. “This achievement not only helps Boxer breeders avoid this disease but it also provides an extraordinary advancement to the study of human heart diseases resulting from electrical conduction defects and the resulting heart muscle changes that occur.”

The disease is well known in Boxers because the breed has the highest incidence of this form of heart disease. ARVC is also known to be an inherited disease and breeders sometimes avoided breeding to certain lines of Boxers yet were never completely sure if those lines had an increased risk of disease. Additionally, the disease tends to vary in severity between different dogs; key indications that the disease had a dominant genetic origin.

Meurs began looking at the disease as an extension of her work with inherited heart disease in cats and dogs. This work is somewhat similar to her work with breeds of cats that also suffer heart disease and for which she has also discovered mutant genes. Her lab developed a molecular probe for these mutations so that cat owners now have a mechanism for screening for the disease and breeding away from it.

Using an extremely powerful gene screening mechanism based on a massive computer chip at the Broad Institute at MIT with investigators Kerstin Lindblad-Toh and Evan Mauceli, Meurs looked at thousands of regions of boxer dogs' DNA simultaneously. The samples were collected with participation by members of the American Boxer Club and the American Boxer Charitable Foundation and were segregated into groups of dogs with the disease and those with no evidence for the disease.

Once computer analysis identified a specific region of interest, Meurs' lab evaluated thousands of DNA sequences in affected and unaffected dogs and identified a gene mutation in a gene that normally codes for the production of a key cellular adhesive protein. Subsequent studies done by WSU veterinary cardiologist, Sunshine Lahmers, demonstrated that the cellular adhesive proteins were located at the junction between cells in the heart.

Theoretically, the conduction defect is in some way responsible for a rapid, irregular heart beat that does not pump blood efficiently. When blood is not pumped efficiently, there may not be enough circulation maintained in the brain and other organs. This can lead to fainting episodes or even sudden cardiac death.

Over time, the right, lower chamber of the heart, called the right ventricle, begins to be infiltrated by a fibrous fatty tissue and often has decreased contractile ability. This change in the heart's tissues can spread to the wall between the heart chambers and even the left ventricle.

The structural changes that result in functional impairment is the hallmark sign seen when a post mortem examination is performed on the animal's heart. Under the microscope, the normal muscle appears solid and dense. The affected heart muscle tissue is riddled with holes where the fibrous fatty tissue has infiltrated stretching it like unorganized lace.

Meurs' laboratory is now near obtaining a patent on her discovery and is perfecting a genetic testing probe for the gene mutation that will be used as a clinical screening device. Shortly, Boxer owners will have the ability to take a simple cheek swab of their dog and know whether or not it carries the mutant gene. Cost of the screening is expected to be about \$70 and available within the next 1-2 months.

"In many cases, after the disease is diagnosed it can be managed with medication for a long enough period of time in a dog's life that other diseases such as cancer will be the cause of death," said Meurs. "The medications are not very expensive and there are generic forms available, too. Average monthly costs are probably less than \$100."

Meurs said that, with her lab's service, Boxer owners and breeders will be able to identify dogs with the mutant gene and are likely to breed away from the disease.

Click [here](#) to see a video from magnetic resonance imaging of both a normal dog's heart and a heart with ARVC.