# Freckle

## on

# Her







## THE

PUG PUG PUG





The Newest Holt Family Member Tugg Pug Pug



# Roo

The Newest Newest Holt Family Member

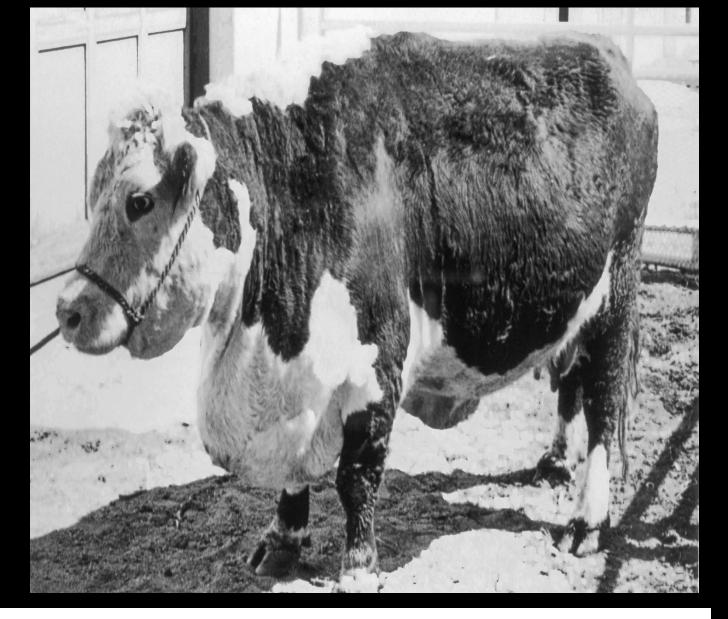


## **Bovine High Mountain Disease**



3000 feet

"Brisket Disease" **Feedlot Cardiac Failure Pulmonary** Hypertension BIF Calgary, Canada Obesity 2023 Coronary Vascular Disease?8000 feet Tim Holt, DVM



#### 1913-First Formal Description of "Dropsy of High Altitude"

PAP Testing at this Elevation

#### 9520 feet

#### 2901.7 meters



### Bovine High Mountain Disease Vs. Feedlot Cardiac Death Alamosa Colorado 8000 feet 3000 Feet

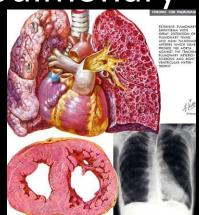






### Background Pulmonary Hypertension Human Aspect

- Hypoxic pulmonary hypertension is the most common cause of cor pulmonale.
- Occurs with alveolar hypoxia, in conditions such as emphysema, sleep apnea. A specialized cause is generalized alveolar hypoxia of altitude, high altitude pulmonary hypertension.
- Early human research



#### PH and the WHO Classification



The bovine model recapitulates important features of Group III PH, the second largest group of PH human patients (Gretta)  Group I pulmonary arterial hypertension, Congenital (PAH)

Group II PH due to left heart disease

Group III PH due to lung diseases and/or chronic hypoxia

Group IV chronic pulmonary thromboembolic PH

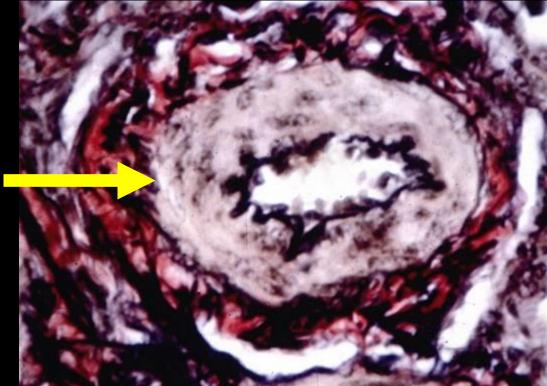
 Group V PH with unclear multifactorial mechanisms

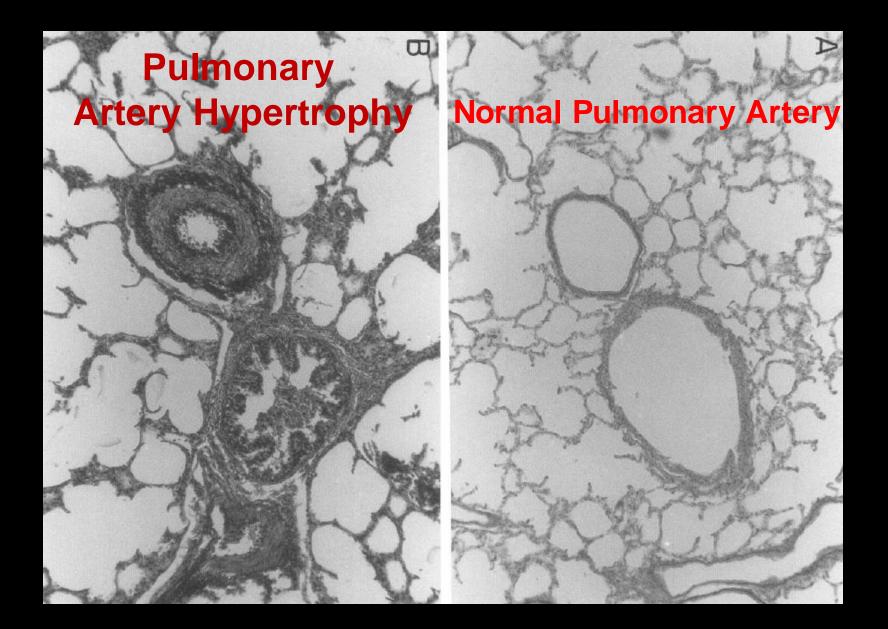
Pathophysiology **Most Critical** BHMD Part Genetics Alveola<u>r</u> Hypoxia **Environmental Pulmonary Vasoconstriction** Don't Pulmonary Remodeling Forget Pulmonary Hypertension Donnie **Right Ventricular Hypertrophy** Another **Right Ventricular Dilation** Genetic Aspect, To **Right Heart Congestive Failure** Die or Not to Die



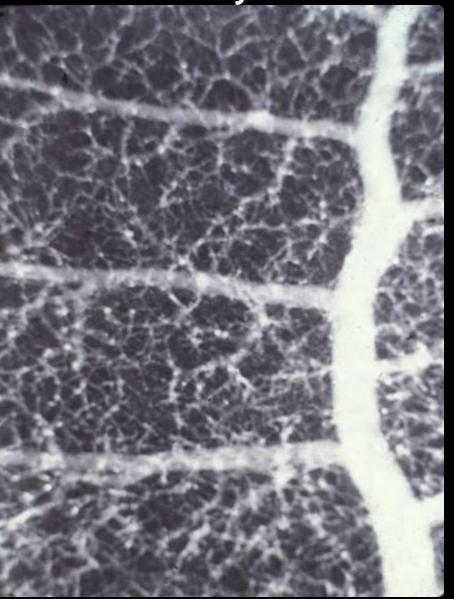
### Normal Pulmonary Artery PAP=40mmHg

Hypertrophy of Pulmonary Artery Smooth Muscle PAP=80mmHg





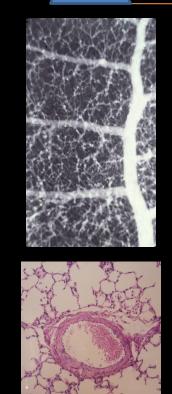
#### 40 mmHg PAP (Normal) Pulmonary Vasculature



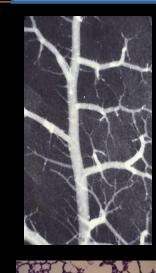
#### 80 mmHg PAP (Vasoconstriction) Pulmonary Hypertension

# Pathophysiology

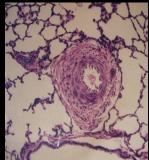
- 1. Alveolar hypoxia Altitude
- 2. Pulmonary arterial vasoconstriction and shunting
- 3. Pulmonary arterial remodeling and arterial smooth muscle hypertrophy



3,200 ft

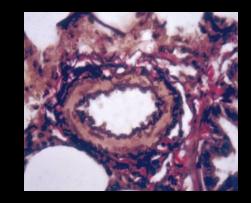


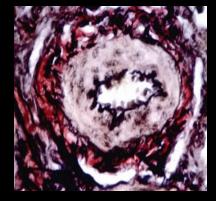
7,240 ft



# Pathophysiology

- 4. Pulmonary hypertension
- 5. Right ventricular hypertrophy
- 6. Right ventricular dilation



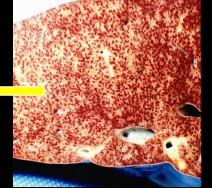




Enlarged Right Ventricle

7. Right congestive heart failure

Congested Liver "Nutmeg Liver"





Clinical Signs Congestive Right Heart Failure BHMD, Feedlot Cardiac Death

-Lethargy, weakness, collapse -Jugular Distention/Pulsation

- -Diarrhea
- -Subcutaneous Edema Brisket Region Intermandibular Ventral Abdomen



-Fluid Within Abdomen Thorax Pericardium

-Bulging eyes

-Death

## Jugular Distension, BHMD 27% Death Loss in One Year 117/435 calves >\$100,000.00 Loss



Wyoming Ranch Elevation 7200 feet Home Ranch

All Calf losses Out of Two Sire Groups

Photo: Thank you, Dr. Knight

## Variation in Response to Hypoxia

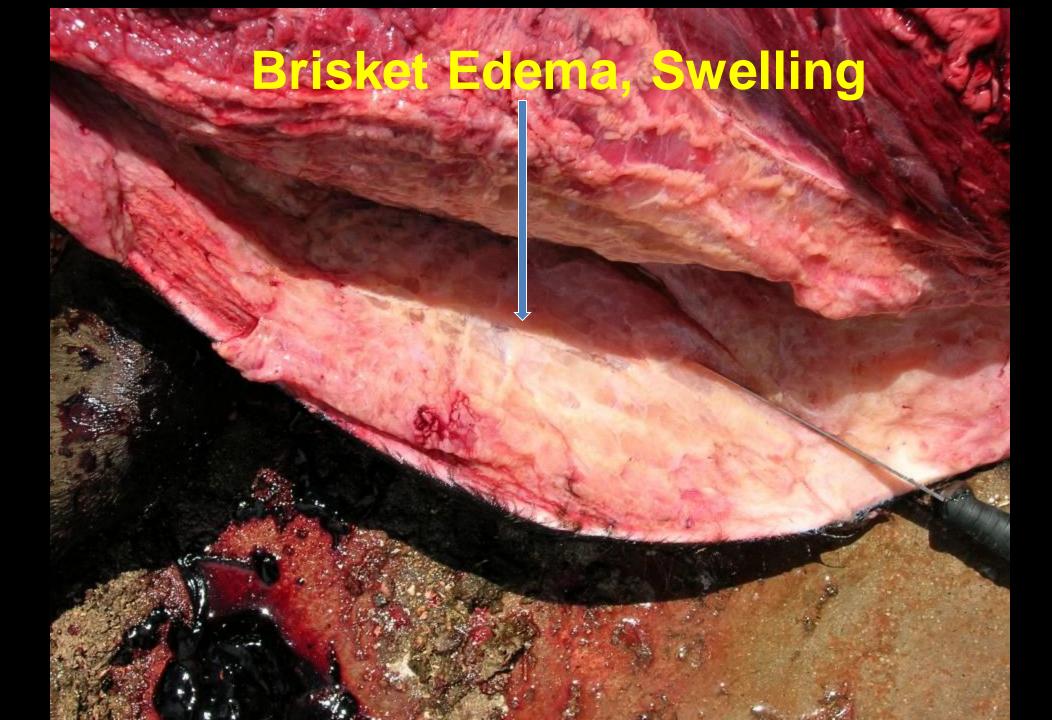
**Hyper-Reactive Bovine**, **Porcine** Moderate-Reactive Equine, (Humans) Hypo-Reactive Ovine, Caprine, Canine, Llama, Alpaca **YAK!!!** 

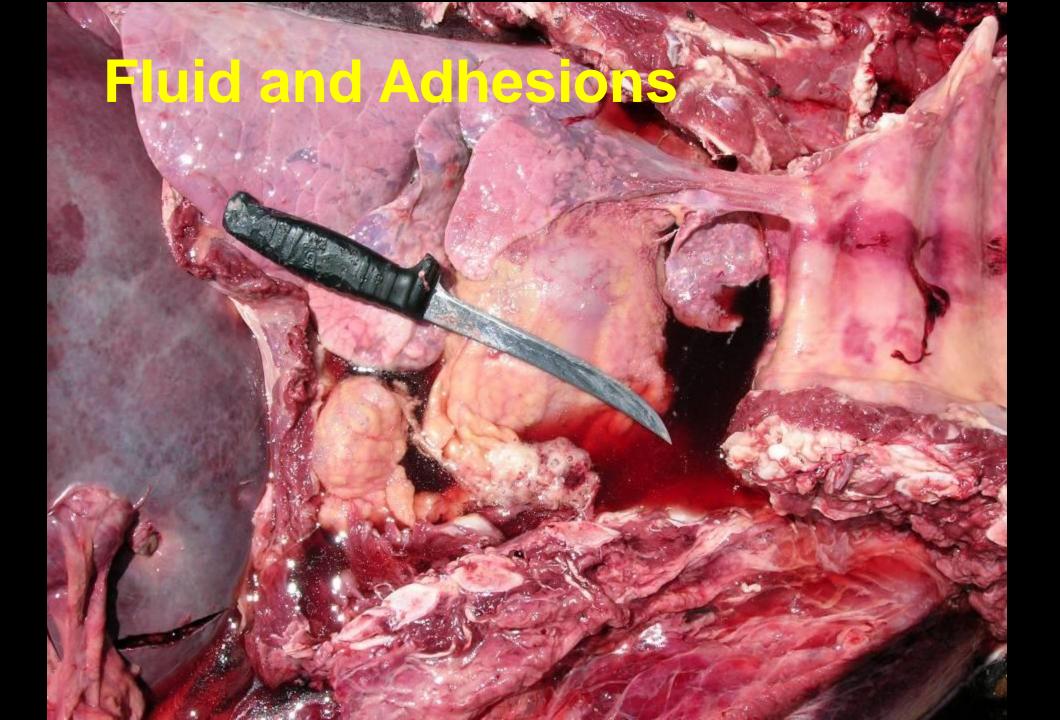
**Breed Susceptibility ?** 





### Necropsy of Clinical Case Of Bovine High Mountain Disease









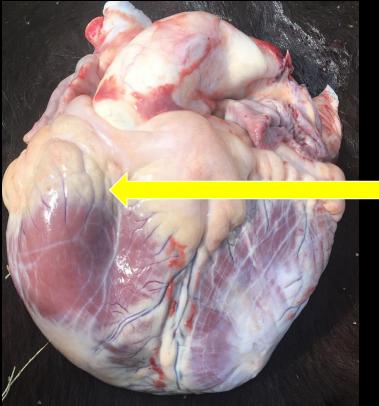
### Aneurism Pending Pulmonary Hypertension





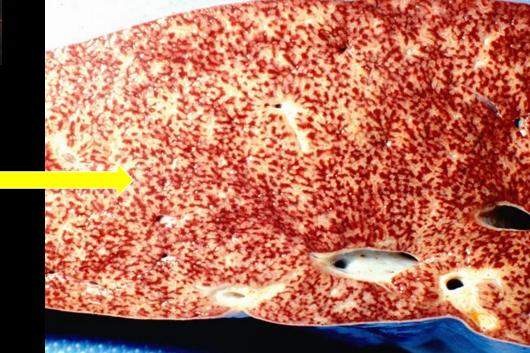
# Congested Liver Swollen Rounded Edges





## Necropsy Findings

Enlarged Right Ventricle GRADE 5 Cardiac Collapse



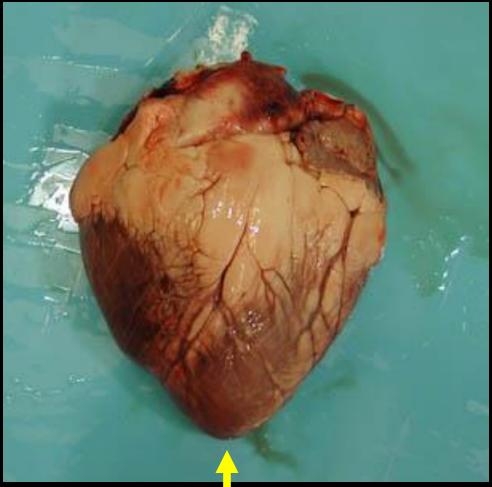
Congested Liver "Nutmeg Liver"

### Dilated Pulmonary Artery

GRADE 5

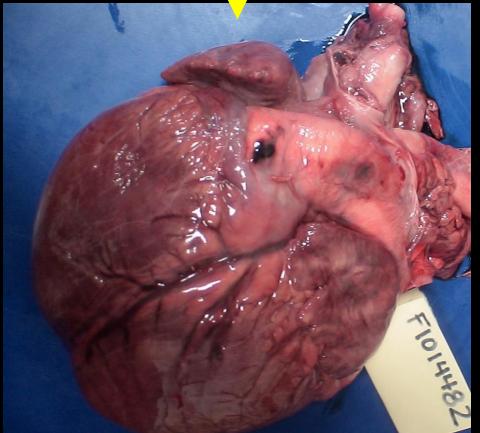
Enlarged Right Ventricle and Atrium

01.13.2009



#### Grade 1—Normal Heart

### Grade 4—Heart, Severe Cardiac Disease





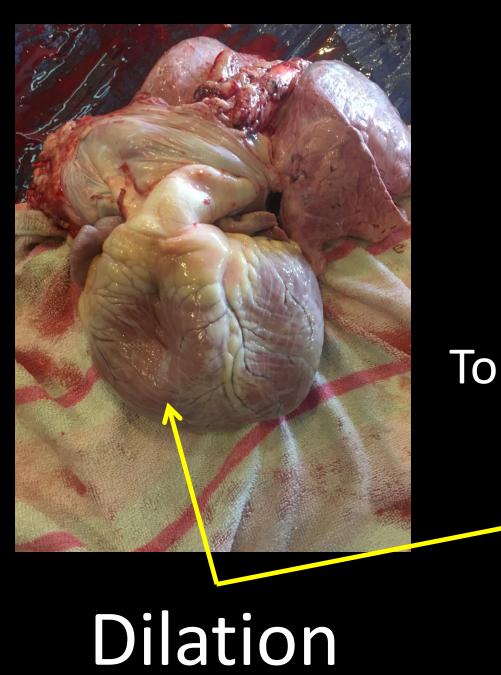
Can we use cardiac anatomical Changes to track Cardiac disease

ABSOLUTELY

### Grade 5 Heart

### Complete Cardiac Collapse





## Hypertrophy









### Why do the Heart Scoring?

Estimated loss of \$250,000.00 per feedlot, incidence continues to rise and is increasing even now

Etiology **genetic driven**, growth, performance area in which to control loss, selection, etc

PAP score has a positive correlation to Higher Heart scores thus Pulmonary Hypertension



### Why do the Heart Scoring

\*Evaluate relationship between healthy heart scores and abnormal heart scores by evaluating phenotypic differences seen in fattening cattle

> \*Better understand the relationship of heart scores in the estimation of heritability for the trait



### Genetics vs. Phenotypic

CSU study in genetics USMARC study, phenotype Simplot Genetics



### High Altitude Disease

- Condition affecting cattle at altitudes of >5,000 ft.
- Pulmonary arterioles begins to constrict and thicken in response to low oxygen being transported.
- Selecting bulls with lower PAP has been successful in producing progeny with lower PAP scores thus more adapted to elevation.

### Feedlot Heart Disease

- Condition affecting feedlot
   cattle at low to moderate
   altitudes.
- Direct cause is currently unknown, but these individuals experience heart remodeling similar to animals experiencing brisket disease.

PAP is currently used as a decision factor for culling animals in the herd who display high PAP at early ages.

HYPOXIA

## **Heart Fat Score**

• \*Epicardial fat has been shown to have relationships with pulmonary hypertension in humans.



### Tracking and Evaluating Incidence of Cardiac Failure Gross Visual External Cardiac Evaluation Grading system 1-5

#### **GRADE: 1, Normal Heart**

-Normal conical shape
-Normal left ventricle apex, easily visible
-Right ventricle smaller than left and in normal portion, fitting on the side of left ventricle
-Normal Atrial Anatomy, right atrium smaller than left
-No clinical evidence of infarction or aneurysm pending, no thinning of vessel wall
-Normal Pulmonary Artery size
-May appear as 2 but with rigor may develop and shrink to 1



#### Gross Visual External Cardiac Evaluation Grading system 1-5

#### **GRADE: 2, Mild Change**

-Normal conical shape

-blunting of left ventricle apex, visible but losing apex point -Right ventricle becoming larger than left, right ventricle becoming pronounced.

-Right Atrium beginning to enlarge, same size or slightly larger than left.

-No clinical evidence of infarction or aneurysm pending, no thinning of vessel wall

 -Pulmonary artery beginning to show mild enlargement.
 -Cardiac muscle when palpated is stiff and suggestive of hypertrophy and loss of lumen space, may be biventricular

-2+, Right Ventricle enlarged but not complete reverse D, flaccid



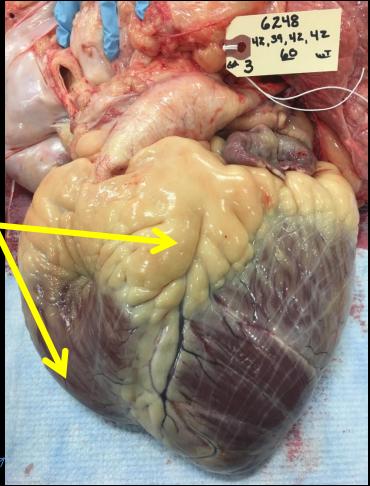
#### Gross Visual External Cardiac Evaluation Grading system 1-5

#### **GRADE: 3, Moderate Change**

Beginning to lose conical shape
Blunting of left ventricle apex, visible but losing apex point and deviating, beginning of reverse "D"
Right ventricle larger than left, right ventricle becoming more pronounced.
Right Atrium enlarged, Atrium larger than left
Can be clinical evidence of infarction or aneurysm

pending, thinning of vessel wall apparent -Pulmonary artery enlarged.

-Cardiac muscle when palpated is stiff and suggestive of hypertrophy and loss of lumen space, may be biventricular, after removal muscle may be severely flaccid



### Gross Visual External Cardiac Evaluation Grading system

**1-5** 

#### **GRADE: 4, Severe Changes**

-Lose of cardiac conical shape
-Left ventricle apex has been lost due to ventricular rounding. Reverse "D" is apparent.
-Right ventricle larger than left, right ventricle becoming more pronounced and taking on rounding shape.
-Right Atrium enlarged, Atrium larger than left and remains congested

-Can be clinical evidence of infarction or aneurysm pending, thinning of vessel wall apparent

-Pulmonary artery enlarged greatly.

-Cardiac muscle when palpated is becoming soft and without shape but still has some muscle tone.



#### Gross Visual External Cardiac Evaluation Grading system 1-5

#### **GRADE: 5, Severe Changes, flaccid heart**

-Loss of cardiac conical shape, severe
-Left ventricle apex has been lost due to
ventricular rounding. Reverse "D" is apparent.
-Right ventricle larger than left, right ventricle becoming
more pronounced and taking on rounding shape.
-Right Atrium enlarged, Atrium larger than left and remains
congested.

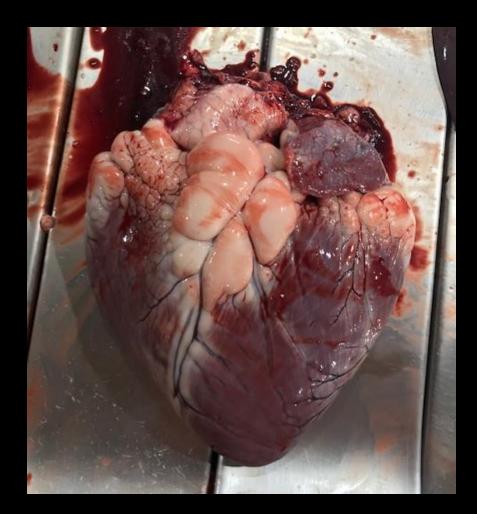
-Can be clinical evidence of infarction or aneurysm pending, thinning of vessel wall apparent

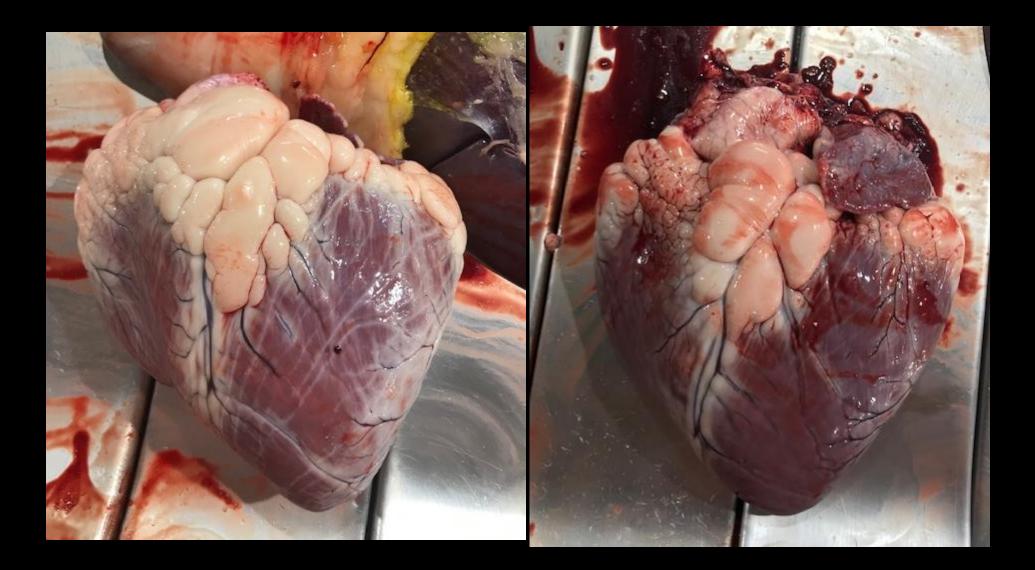
-Pulmonary artery enlarged greatly.

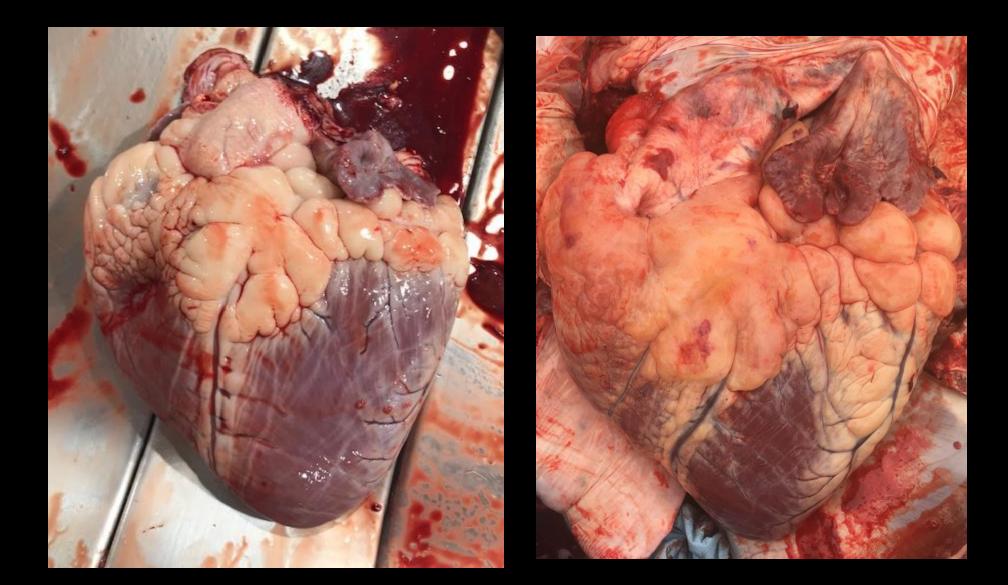
-Cardiac muscle when palpated is soft and without shape, there is no muscle tone and heart lays flat.



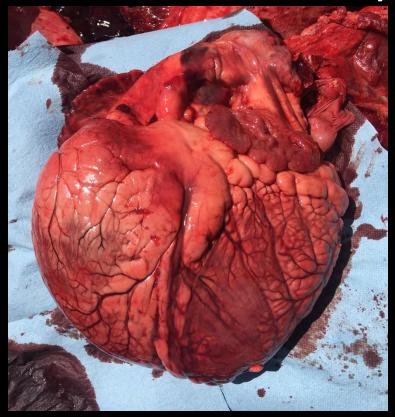




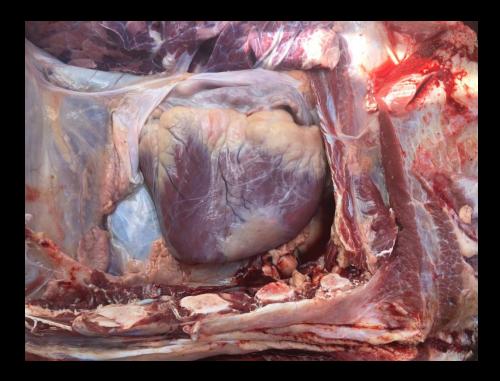














## Heart Scores Summary Thus far

Results thus far:

\*High Heart Scores correlate with High Heart Fat Scores



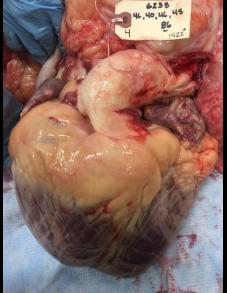
\*There is a high correlation to sire groups

### Conclusions:

Significant score effect, 56 sires accounted for

66%; scored 1,2 normal

34%; 3 plus, significant cardiac remodeling



# **PAP Testing Adventures**

- 1980 in Gunnison/Hesperus Colorado
- 1980-Present,
   >492,000 head
- Ambient Temps -42 degrees to 115 degrees.
- Numerous Breeds with high Pap's in all
- Elevations from sea level to 14,300 feet



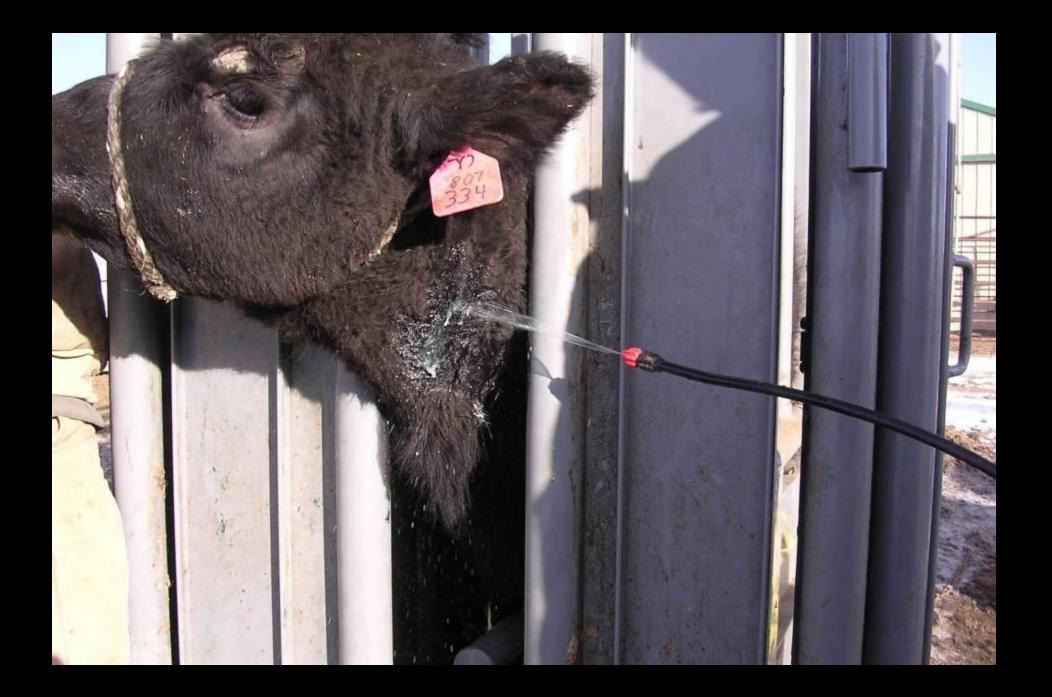














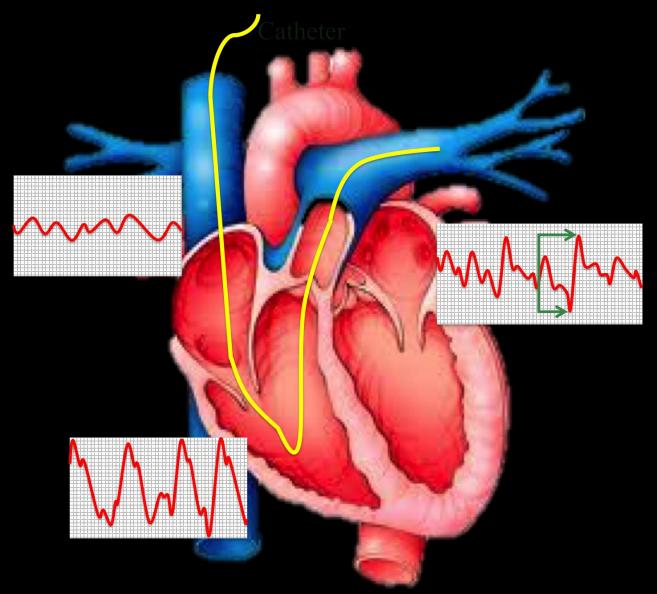




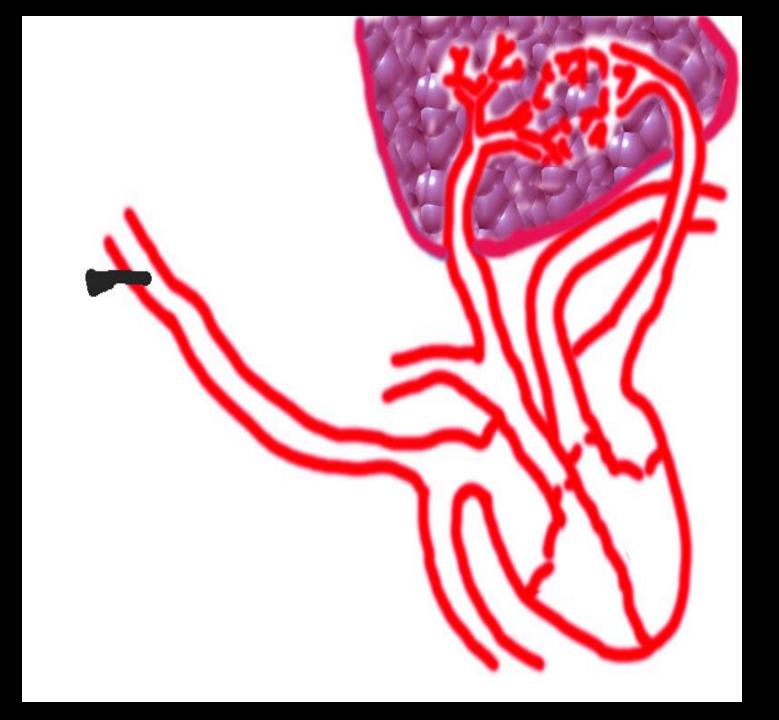


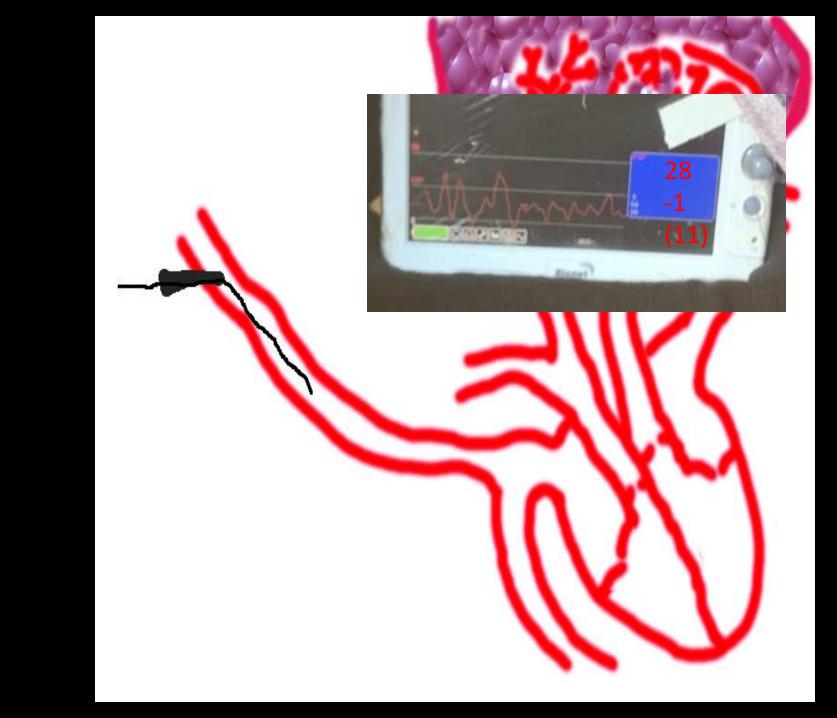


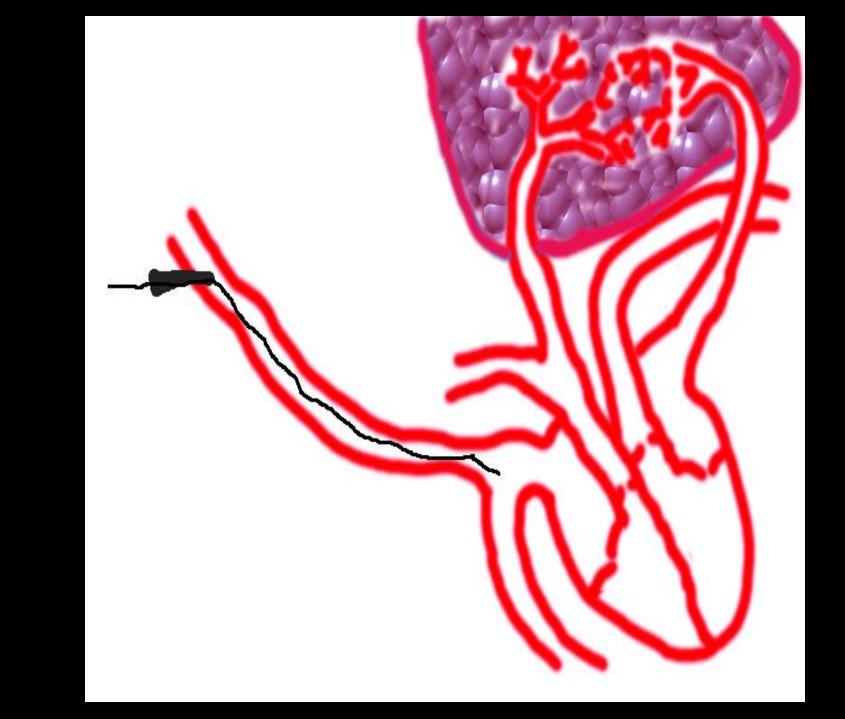
## PAP Measurement Catheter Location

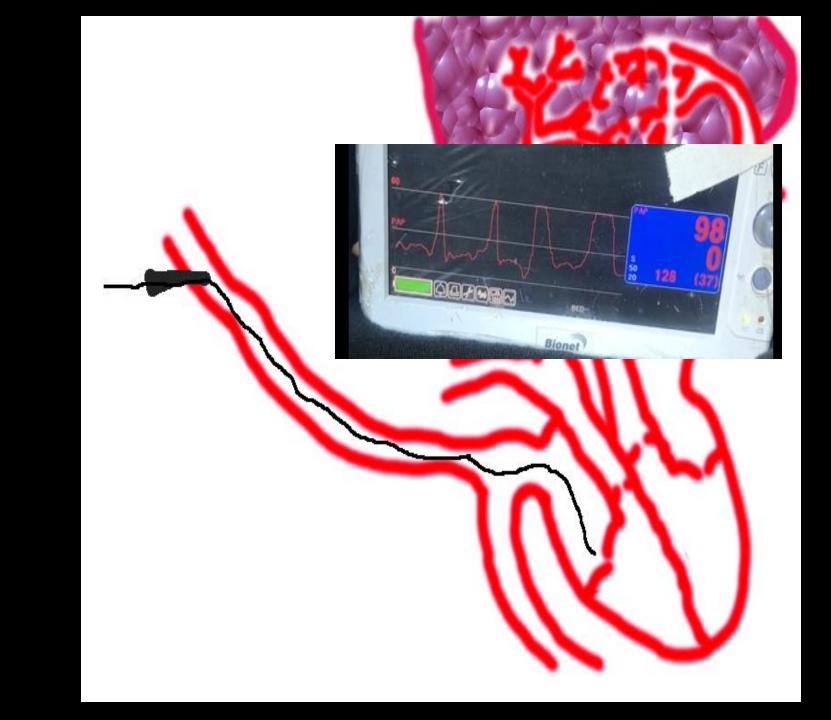


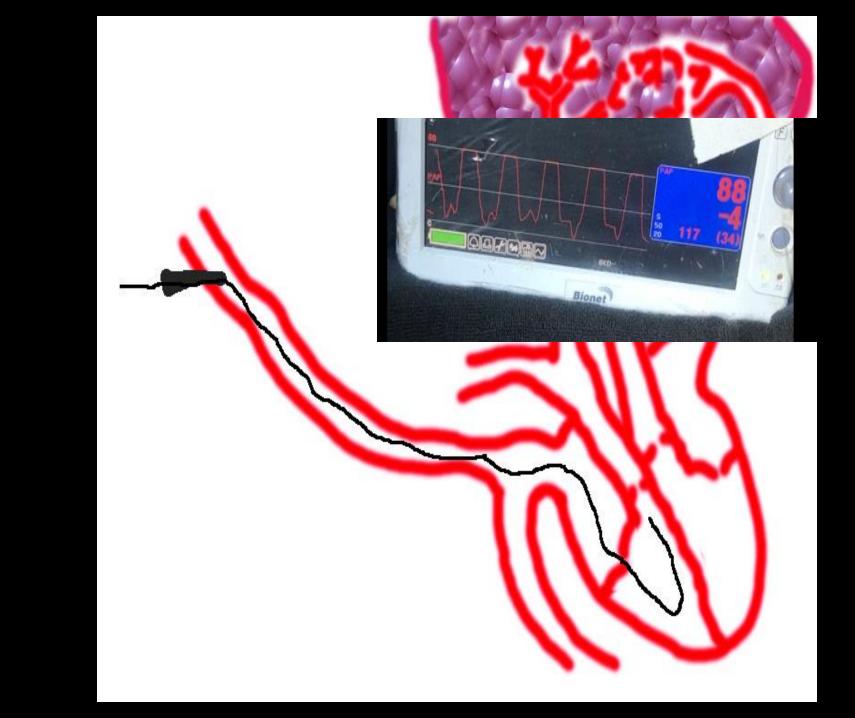


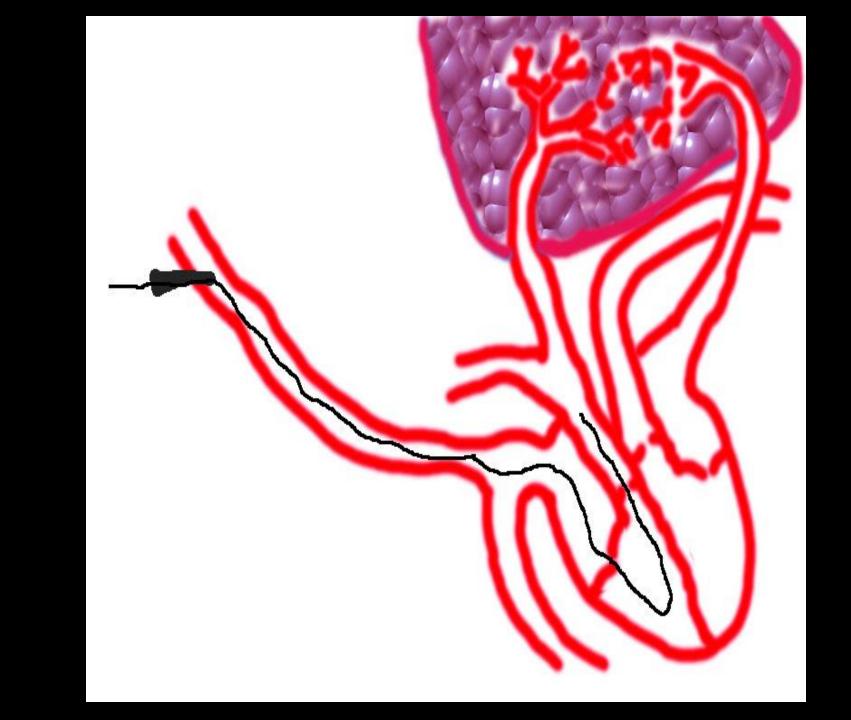


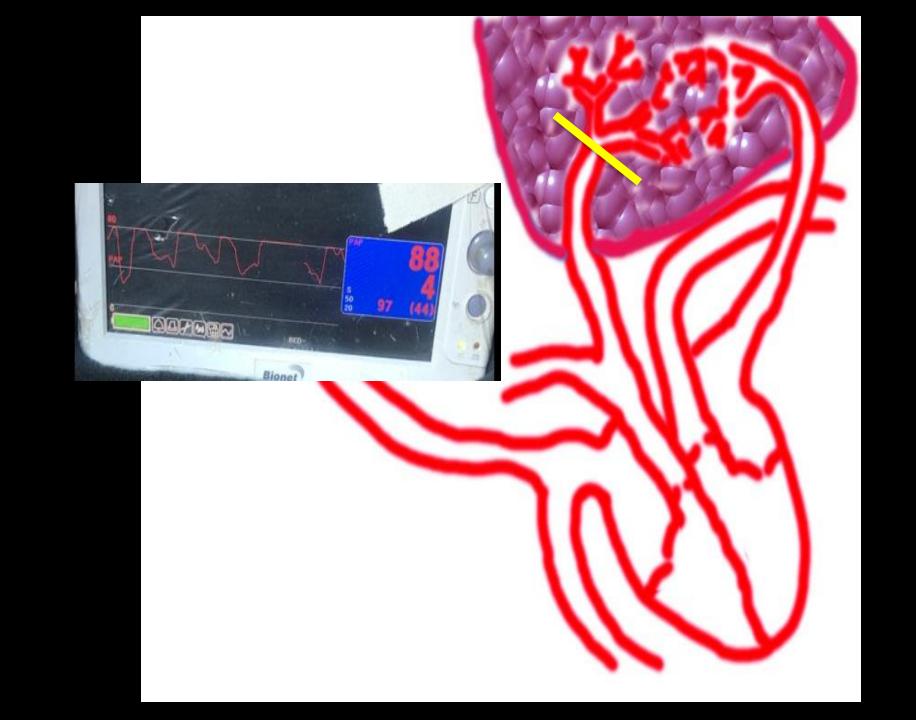




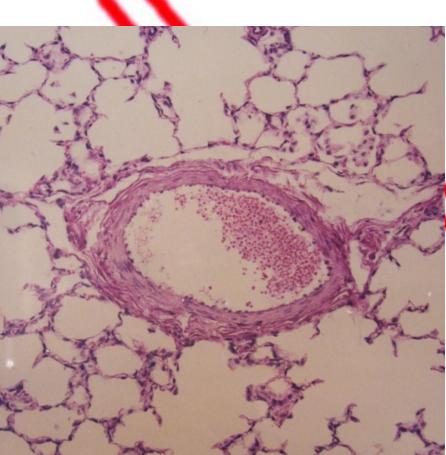








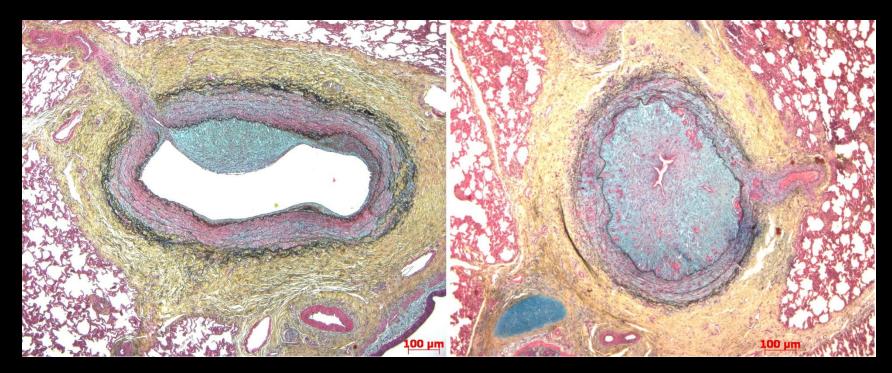








# PAP PAP of of of 46 mmHg 94 mmHg



# Confounding Factors "Body Condition"

- Excessive body condition increases PAP
  - Bulls on Test

 $\bigcirc$ 

- Feedlot Cattle, Newest area of Research!

per year concerning

1.4 billon dollar loss

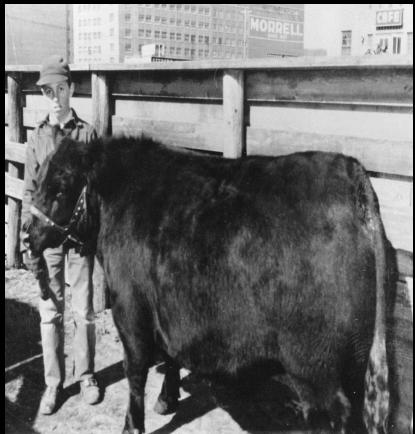
Feedlot

Cardiac Death



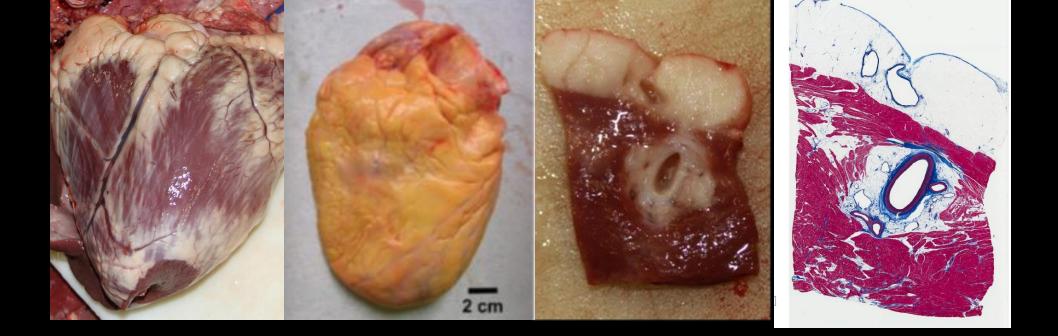
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## Average Live Market Weight has Increased 48% since 1944





1953 Angus Steer 950 lb. Live Weight 2015 Angus Steer 1200-1600 lb. Live Weight





Closer Look at Feedlot Cardiac Death Is It Bovine High Mountain Disease

#### Feedlot Cardiac Death

Fattening Induces Hypercholesterolemia and Dyslipidemia



# **Altitude and Repeatability**

• <5000 feet

Estimated Repeatability 60% Predictive Screening only

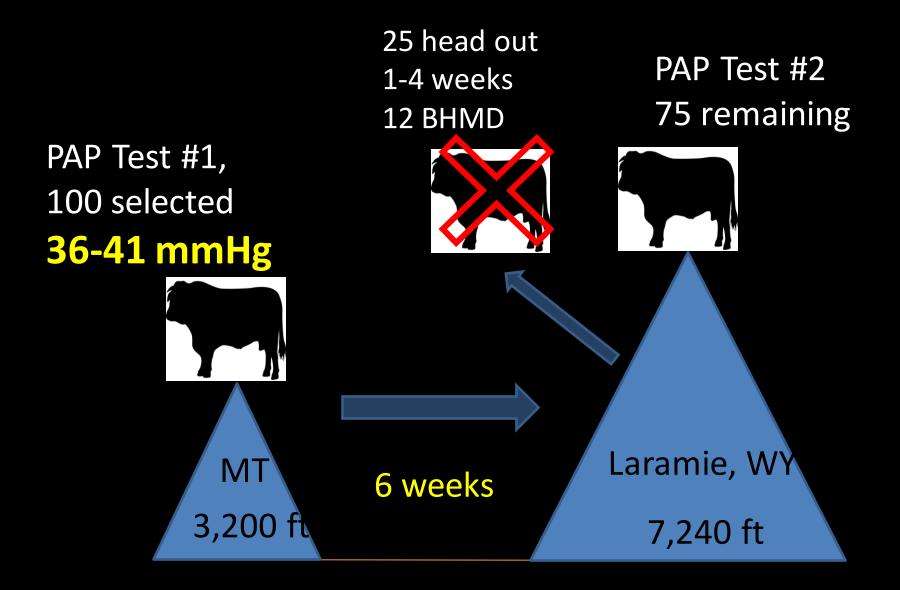


 1-1.5 mmHg increase per 1000 feet elevation rise.

# **PAP Hot Topics**

Elevated PAP—Accurate Measurements <41 Questionable predictive value

## Herd Example: Red Angus Bulls 16 months of Age



# 

- 48% (36/75) had PAPs <a>>46 mmHg</a>
  - -10/75 had PAPs 46-50 mmHg
    - Pulled from sale
  - **-26/75** had PAPs > 50 mmHg
    - Pulled from sale and moved to lower elevation
- 52% (39/75) had PAP <45mmHg
- Average increase of +12 mmHg

#### PAP Risk Factor Low Elevation Test Chart PAP test conducted at elevation <4000 ft. (60% Repeatable, Predictive Value)

PAP	Use at Low Elev.	Use at Moderate Elev.	Use at High Elev.	Use at Extreme
<u>Score</u>	(<4000 feet)	(4000-5500 FEET)	(5500-7500 feet)	<u>(&gt;7500 feet)</u>
34-39	Low Risk	Low Risk	<b>Moderate Risk</b>	High Risk
40-45	Low Risk	<b>Moderate Risk</b>	High Risk	High Risk
<b>46-49</b>	<b>Moderate Risk</b>	High Risk	Do Not Use	Do Not Use
<u>&gt;</u> 50	<b>Moderate Risk</b>	High Risk	Do Not Use	Do Not Use

#### PAP Risk Factor Moderate Elevation Test Chart PAP test conducted at elevation <u>4000-5500 ft.</u> (70% Repeatable, Predictive Value)

PAP	Use at Low Elev.	Use at Moderate Elev.	Use at High Elev.	Use at Extreme
<u>Score</u>	(<4000 feet)	(4000-5500 FEET)	(5500-7500 feet)	<u>(&gt;7500 feet)</u>
34-39	Low Risk	Low Risk	Low Risk	Moderate Risk
40-45	Low Risk	Low Risk	<b>Moderate Risk</b>	High Risk
<b>46-49</b>	Moderate Risk	High Risk	Do Not Use	<b>Do Not Use</b>
<u>&gt;</u> 50	Moderate Risk	High Risk	Do Not Use	Do Not Use

#### PAP Risk Factor High Elevation Test Chart PAP test conducted at elevation <u>5500-7000 ft.</u> (75-95% Repeatable, Predictive Value)

PAP	Use at Low Elev.	Use at Moderate Elev.	Use at High Elev.	Use at Extreme
<u>Score</u>	(<4000 feet)	(4000-5500 FEET)	(5500-7500 feet)	<u>(&gt;7500 feet)</u>
34-39	Low Risk	Low Risk	Low Risk	Low Risk
40-45	Low Risk	Low Risk	Moderate Risk	<b>Moderate Risk</b>
<b>46-4</b> 9	Moderate Risk	<b>Moderate Risk</b>	<b>Moderate Risk</b>	High Risk
<u>&gt;</u> 50	<b>Moderate Risk</b>	<b>Moderate Risk</b>	High Risk	High Risk

#### **PAP Risk Factor**

### Very High Elevation Test Chart PAP test conducted at elevation >7000 ft. (95% Repeatable, Predictive Value)

PAP	Use at Low Elev.	Use at Moderate Elev.	Use at High Elev.	Use at Extreme
<u>Score</u>	(<4000 feet)	(4000-5500 FEET)	(5500-7500 feet)	<u>(&gt;7500 feet)</u>
<b>34-39</b>	Low Risk	Low Risk	Low Risk	Low Risk
<b>40-45</b>	Low Risk	Low Risk	Low Risk	<b>Moderate Risk</b>
<b>46-49</b>	<b>Moderate Risk</b>	<b>Moderate Risk</b>	<b>Moderate Risk</b>	High Risk
<u>&gt;</u> 50	<b>Moderate Risk</b>	<b>Moderate Risk</b>	High Risk	High Risk

### WHY ETHIOPIA?

- Ethiopian highlanders maintain venous hemoglobin concentrations and arterial oxygen saturation within the ranges of sea level populations, despite the decrease in the ambient oxygen tension at high altitude.
- Different from Andean and Tibetan populations







# PAP testing in Ethiopia Elevation 14, 300 feet

Tested 325 Head Highest PAP Measurement 34 mmHg



# Questions



# Finished



# Thank You !!

# Questions



# Finished



# Thank You !!