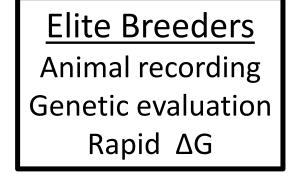
### Animal Breeding Systems for Small Ruminants David Notter Dept. of Animal & Poultry Sciences Virginia Tech









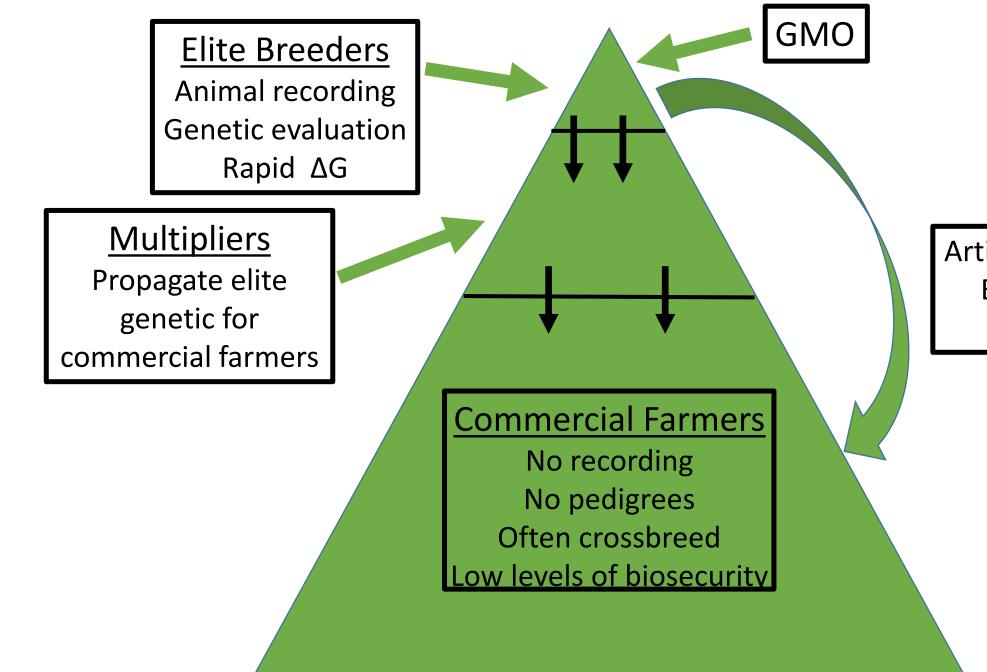


**Multipliers** 

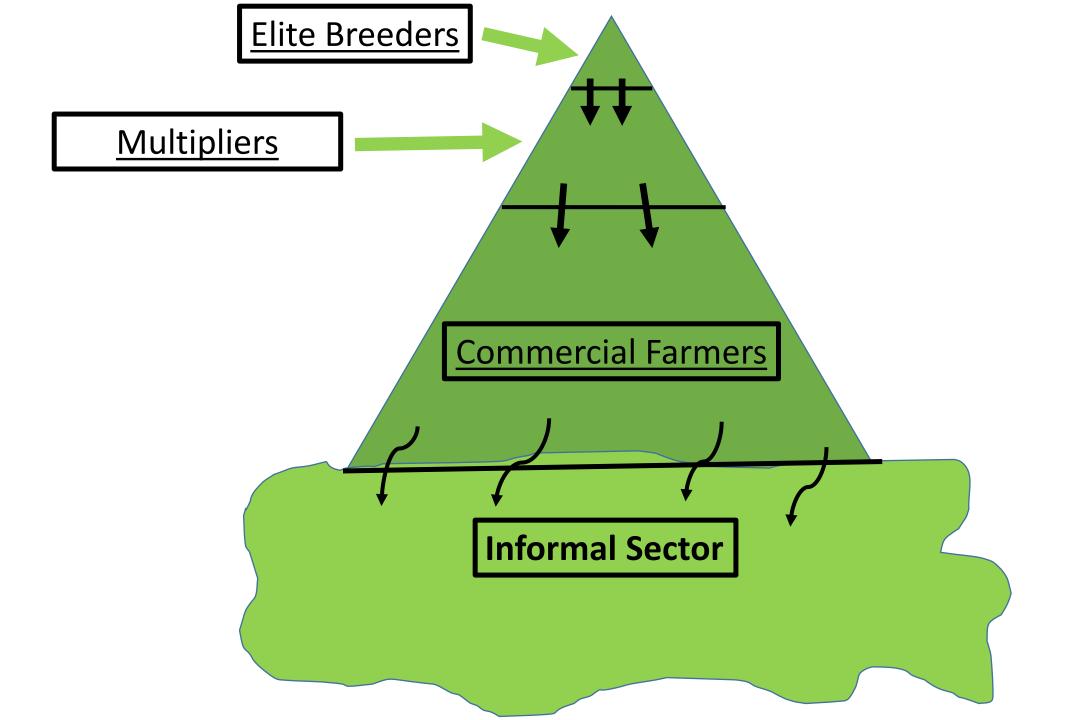
Propagate elite genetic for commercial farmers

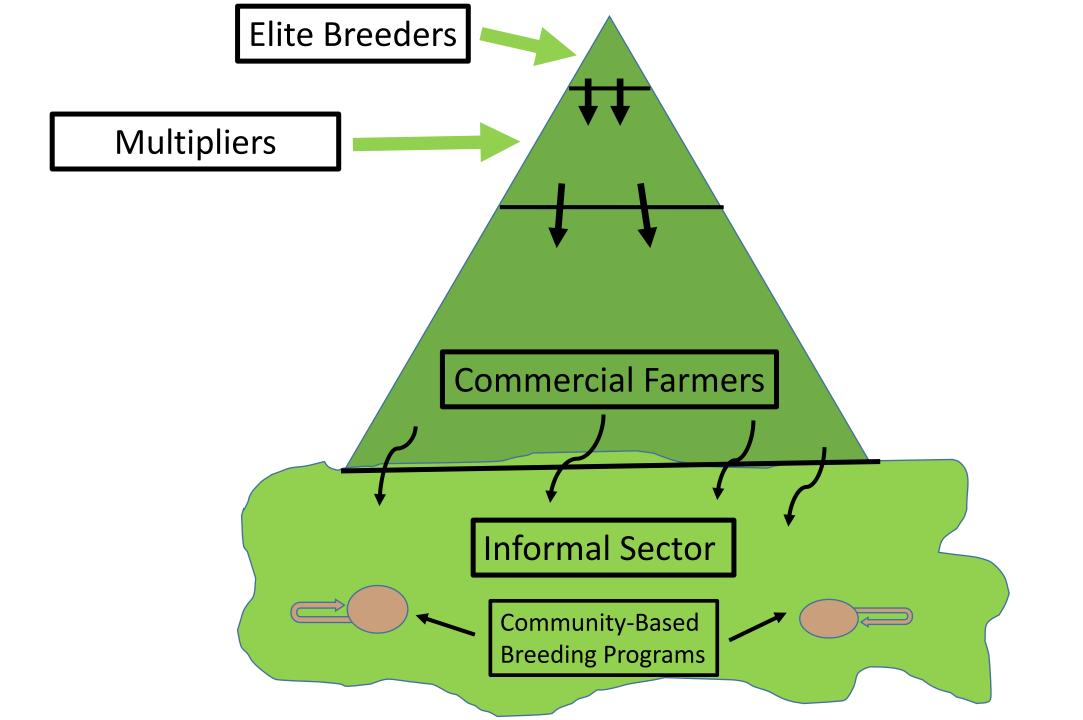
**Commercial Farmers** 

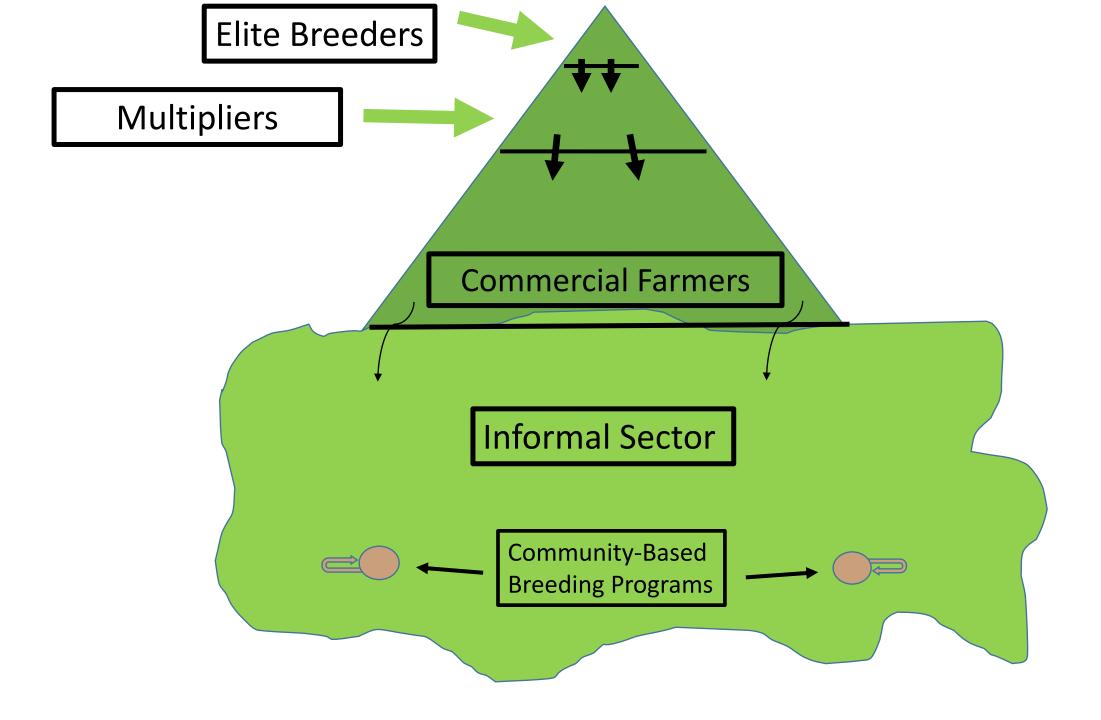
No recording No pedigrees Often crossbreed Low levels of biosecurity



Artificial Insemination Embryo Transfer Cloning







Candidates for Gene Manipulation in Small Ruminants—Production Traits

- Litter size: objective is to increase the frequency of twins without producing litters of 3 or more
- Muscularity: increase muscle growth to enhance lean meat yield
- Fiber (cashmere, wool) production: increase fiber production without also increasing fiber diameters (fineness)
- Color: modify color inheritance to facilitate management of crossbreeding programs
- A2 milk (cattle) to take advantage of perceived health benefits

# Candidates for Gene Manipulation in Small Ruminants Production Traits

- Most of these changes involve a loss in regulatory control, or an effective "knock-out" of an otherwise functional gene that, under past conditions, had selective value for the species.
- This is easier than doing a precise base substitution, as any (most?) disruptions will do.
- But multiple disruptions are possible, creating regulatory challenges, even if effects on the animal are identical.
- Losses of regulatory control often require compensating increases in environmental control to take advantage of the enhanced performance potential.

#### Candidates for Gene Manipulation in Small Ruminants Production Traits Litter Size

- Litter size: objective is to increase the frequency of twins without producing litters of 3 or more.
- Loss-of-function mutations in several genes can increase ovulation rates and, potentially, litter size in sheep, with likely homologues in goats.
- Candidate target genes include BMPR-1B (Booroola), BMP-15 (sex-linked), GDF-9 (multiple mutations). Many of these mutations can result in litters of 3 to 5 lambs, which is not desirable.
- Homozygotes for some, but not all, mutations in BMP-15 and GDF-9 are sterile, so careful breeding management is required for utilization. New GDF-9 mutation from Brazil has both a modest effect and no homozygote sterility.
- Garole Deccani story

# Candidates for Gene Manipulation in Small Ruminants Production Traits Muscularity

- Primary candidate for manipulation is myostatin, which normally regulates muscle growth.
- Several loss-of-function mutations in cattle produce the "double-muscled" condition. Similar mutations exist in sheep and goats, but generally with smaller effects and possible effects of genetic background.
- Potential for more intensive study of myostatin sequences in sheep and goats to identify better candidates for manipulation.
- Advantages can only be captured if the nutrition of the animal is adequate.
- Callipyge story.



#### Texel

Found in the Netherlands & Belgium
Possesses a mutation in the regulatory region of the myostatin gene that results in partial loss of normal regulation of muscle growth





# Candidates for Gene Manipulation in Small Ruminants Production Traits Fiber Production

- Primary candidate for manipulation is fibroblast growth factor 5 (FGF-5), which normally regulates hair growth.
- Naturally occurring mutations exist in several species.
- Studies in China have documented greater growth of both hair and cashmere in young cashmere goats, without compensating increases in fiber diameter.
- Potential value will depend on the interactions of fiber weight and fiber quality in defining value, e.g., Australian Merino.
- Increasing the length of the cashmere does have economic value.
- Potential homologues in alpaca?

#### Candidates for Gene Manipulation in Small Ruminants Production Traits Color

- Crossbreeding is often problematical in developing countries because of inadequate control over mating plus the tendency to retain attractive (i.e., big) but poorly adapted individuals as breeding females.
- Depending on the color of the local females, a sire breed for crossing could be developed with a color marker (e.g., dominant black in sheep) that would mark the crossbred offspring.

#### A2 Milk

- A2 variant of the beta-casein milk protein.
- Potential to create A2 milk lines from elite Holstein cattle.

Both of these would require precise, targeted manipulations, not just loss-offunction.

# Potential for Gene Manipulation in Small Ruminants Health and Disease

- Disease resistance/resilience can be complex and multifactorial (gastrointestial parasites; trypanosomiasis).
- Manipulation of immune function can have unanticipated effects.
- Limited biosecurity in pastoral production.
- Can disease-resistant GM animals become a reservoir of disease for non-GM individuals or species?
  - Pasteurellosis in Bighorn sheep, avian influenza in wild waterfowl?
- Does resistance to one disease impair resistance to others?
  - Ovine progressive pneumonia/Maedi-visna?
- Scrapie story.
- Regulators need independent input from both specialists and generalists

## Conclusions Regulatory Issues in Evaluating GM Animals

- Human health risks
  - Modest (very modest?) for production traits
  - Avoid creating vectors for potentially zoonotic diseases
- Environmental impacts
  - Unanticipated impacts from use of GM and gene drive to dramatically reduce populations of disease vectors (mosquito, Tsetse fly, ticks, etc.).
  - Domestic animals as reservoirs of diseases for wild relatives?
- Animal health and welfare
  - Direct effects on animal welfare: genetic sterility, reduced resilience to certain diseases, increased frequency of embryo loss or congenital genetic defects (all highly speculative)
  - Indirect effects on animal welfare through inadequate nutrition or management.