

**Understanding contagious transmission of *Mannheimia haemolytica* in feedlot calves by leveraging whole genome sequencing of a unique isolate collection**



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**OBJECTIVE**

The goal of this project is to evaluate the contagious spread of *Mannheimia haemolytica* (*Mh*) between cattle in a feedlot setting, by utilizing whole-genome sequencing to analyze genetic relationships of *Mh* between contiguous pens of cattle.

**METHODS**

- Deep nasopharyngeal swabs were collected at arrival, 13 DOF, and 36 DOF from 4 pens of fall 2020 placed calves in a Saskatchewan research feedlot, with 100 calves per pen (400 calves total)
- Cattle received a 5-way BRD vaccine with *Mh* toxoid, clostridial vaccine, and metaphylaxis with tulathromycin
- All calves were sampled at Day 0 and Day 13, and 10 calves per pen were sampled at Day 36
- 489 total *Mh* isolates were found on culture
- Whole genome sequencing was performed on all isolates with Illumina NovaSeq 6000 SP PE150
- Genomes were assembled with Shovill and annotated with Prokka 1.14.6. Resistance genes were identified using ABRicate.
- Alignments were constructed with CSI Phylogeny, using USMARC strain 191 as a reference, to identify genetic relationships between isolates.

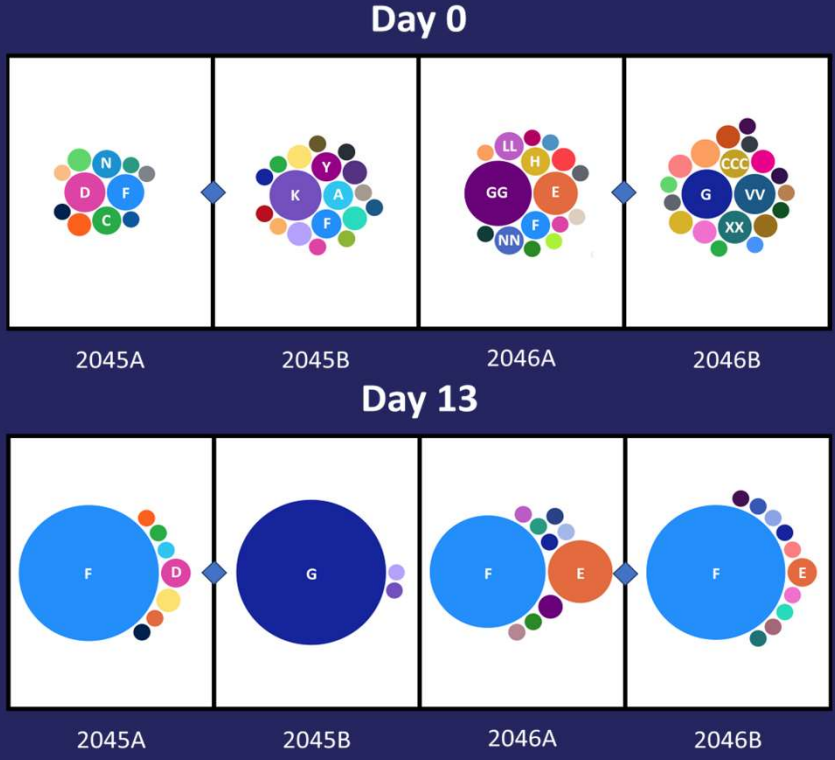
**RESULTS**

- At arrival there were a total of 154 isolates, spread between 56 different clusters.
- At 13 DOF, there were a total of 307 different isolates, spread between 26 different clusters.
- In each pen, one cluster became dominant by 13 DOF.
- At 36 DOF, only 10 calves per pen were sampled. Total number of *Mh* isolates was 28, in 10 clusters.
- There were 8 different resistance genes identified. Classes include aminoglycosides/cyclitols, macrolides, sulfonamides, and tetracyclines
- Clusters F and G had a "resistance profile" characterized by genes *mphE*, *msrE*, and *sul2*
- Clusters E and LL had a "resistance profile" characterized by genes *strA*, *aphA1*, *strB*, *estT*, *sul2*, and *tetH*

**CONCLUSION**

This study provided additional evidence to show that selection for and expansion of a dominant strain of *Mh* appears to be a not uncommon occurrence in cattle entering feeding operations.

# A single clone of *Mannheimia haemolytica* becomes dominant within a pen just 13 days after arrival.



Blue diamond indicates a shared water source

Fig. 1 and 2. Cluster distribution of *Mannheimia haemolytica* isolates collected from all cattle at 0 and 13 DOF.

Table 1. Calves shedding *Mannheimia haemolytica* as identified by whole-genome sequencing, by pen and sampling time. Each pen contains 100 calves in total.

| Pen   | Time Point<br>No. isolates (%) |         |        |
|-------|--------------------------------|---------|--------|
|       | 1 DOF                          | 13 DOF  | 36 DOF |
| 2045A | 26 (26)                        | 77 (77) | 8 (80) |
| 2045B | 36 (36)                        | 83 (83) | 7 (70) |
| 2046A | 46 (46)                        | 68 (68) | 6 (60) |
| 2046B | 46 (46)                        | 79 (79) | 7 (70) |

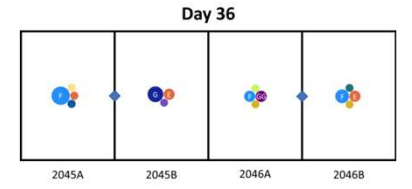


Fig. 3. Cluster distribution of *Mannheimia haemolytica* isolates collected from 10 cattle per pen at 36 DOF.

Table 6. Resistance gene distribution by cluster within pens at 0 DOF. Clusters without resistance genes are not shown. No resistance genes were identified from isolates from pen 2046B on day 0.

| Class           | Gene         | Pen (Total No. Isolates) |       |            |       |            |       |
|-----------------|--------------|--------------------------|-------|------------|-------|------------|-------|
|                 |              | 2045A (26)               |       | 2045B (36) |       | 2046A (46) |       |
|                 |              | D (6)                    | F (5) | F (3)      | E (7) | F (3)      | LL(3) |
| Aminoglycosides | <i>strA</i>  | 0                        | 0     | 0          | 7     | 0          | 3     |
|                 | <i>aphA1</i> | 0                        | 0     | 0          | 7     | 0          | 3     |
|                 | <i>strB</i>  | 0                        | 0     | 0          | 7     | 0          | 3     |
| Macrolides      | <i>mphE</i>  | 1                        | 5     | 3          | 0     | 3          | 0     |
|                 | <i>msrE</i>  | 1                        | 5     | 3          | 0     | 3          | 0     |
|                 | <i>estT</i>  | 0                        | 0     | 0          | 7     | 0          | 3     |
| Sulfonamides    | <i>sul2</i>  | 1                        | 5     | 3          | 7     | 3          | 3     |
| Tetracyclines   | <i>tetH</i>  | 0                        | 0     | 0          | 7     | 0          | 3     |

Table 7. Resistance gene distribution by cluster within pens at 13 DOF. Clusters without resistance genes are not shown.

| Class           | Gene         | Pen (Total No. Isolates) |        |            |        |            |       |
|-----------------|--------------|--------------------------|--------|------------|--------|------------|-------|
|                 |              | 2045A (77)               |        | 2045B (82) |        | 2046B (79) |       |
|                 |              | E (1)                    | F (67) | G (80)     | F (44) | LL(1)      | E (3) |
| Aminoglycosides | <i>strA</i>  | 1                        | 0      | 0          | 13     | 0          | 1     |
|                 | <i>aphA1</i> | 1                        | 0      | 0          | 13     | 0          | 1     |
|                 | <i>strB</i>  | 1                        | 0      | 0          | 13     | 0          | 1     |
| Macrolides      | <i>mphE</i>  | 0                        | 67     | 80         | 1      | 44         | 0     |
|                 | <i>msrE</i>  | 0                        | 67     | 80         | 1      | 44         | 0     |
|                 | <i>estT</i>  | 1                        | 0      | 0          | 14     | 0          | 1     |
| Sulfonamides    | <i>sul2</i>  | 1                        | 67     | 80         | 14     | 44         | 1     |
| Tetracyclines   | <i>tetH</i>  | 1                        | 0      | 0          | 13     | 0          | 1     |

