The effects of administering different metaphylactic antimicrobials on growth performance and health outcomes of high-risk, newly received feedlot steers

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INTRODUCTION

Bovine respiratory disease (BRD) is the primary animal health concern facing feedlot producers. Many antimicrobial mitigation strategies are available, but few studies have compared feedlot performance during both the receiving and finishing periods following application of different antimicrobials used as metaphylaxis at arrival. The objective of this study was to compare antimicrobial metaphylaxis methods on clinical health and growth performance across both the receiving and finishing periods. A total of 238 multiple-sourced steers in two source blocks were used in a generalized complete block design. The four treatments included: 1) a negative control, 5 mL of sterile saline injected subcutaneously (CON); 2) subcutaneous administration of florfenicol at 40 mg/kg of BW (NUF); 3) subcutaneous administration of ceftiofur in the posterior aspect of the ear at 6.6 mg/kg of BW (EXC); and 4) subcutaneous administration of tulathromycin at 2.5 mg/kg of BW (DRA). The morbidity rate for the first treatment of BRD was decreased for the DRA and EXC treatments compared to CON and NUF ($P < 0.01$). Additionally, average daily gain (ADG), dry matter intake (DMI), and gain-to-feed (G:F) were greater ($P < 0.02$) in the DRA treatment during the receiving period compared to all other treatments. The ADG was also greater ($P < 0.05$) for EXC than the CON treatment throughout the finishing period. Nonetheless, other growth performance variables did not differ among metaphylactic treatments during the finishing period ($P \geq 0.14$). Likewise, no differences in carcass characteristics or liver abscess score were observed ($P \geq 0.18$). All complete blood count (CBC) variables were affected by day ($P \leq 0.01$) except mean corpuscular hemoglobin concentration ($P = 0.29$). Treatment × time interactions were observed for platelet count, white blood cell (WBC) count, monocyte count and percentage, and lymphocyte percentage ($P \leq 0.03$). However, there were no observed hematological variables that differed among treatment ($P \geq 0.10$). The results indicate that some commercially available antimicrobials labeled for metaphylactic use are more efficacious than others in decreasing morbidity rate.

Key words: beef cattle, complete blood counts, growth performance, health outcomes, metaphylaxis

MATERIALS AND METHODS

Experimental procedures were approved by the Texas Tech University Institutional Animal Care and Use Committee.
(approval number 20039-04). The experiment was conducted from October 2020 to August 2021 at the Texas Tech University Burnett Center. The average observed temperature during the study period was 15.1 °C with a maximum of 42.2 °C and minimum of −17.7 °C. The total precipitation was 499.5 mm and average relative humidity was 44%. A chronology of key events (body weight [BW] measurement and blood collection) between blocks 1 and 2 is reported in Table 1.

**Arrival Procedures**

Crossbred steers (N = 238; arrival BW = 248 ± 9.5 kg) were sourced from multiple areas. Steers were blocked by arrival date with block 1 consisting of 123 steers purchased from an auction market in Dalhart, TX and shipped approximately 322 km arriving on October 22, 2020. Block 2 consisted of 115 steers purchased from an auction market in West Plains, MO and shipped approximately 1186 km arriving on December 2, 2020. After arrival, animals were placed into soil surface pens and allowed ad libitum access to water and long-stem grass hay and fed a receiving diet of approximately 65% concentrate at 1% of body weight (BW; Table 2). Within 24 h of arrival (day -1), steers were given an identification tag, individually weighed in a hydraulic squeeze chute calibrated with 454 kg of certified ad libitum access to water and long-stem grass hay and fed a receiving diet of approximately 65% concentrate at 1% of body weight (BW; Table 2). Within 24 h of arrival (day -1), steers were given an identification tag, individually weighed in a hydraulic squeeze chute calibrated with 454 kg of certified weigh cells before weighing with an accuracy of ±0.91 kg (Silencer, Moly Manufacturing, Lorraine, KS), and any bulls (N = 2) were identified, castrated, and not allocated to treatments since they could not be equally stratified among the four treatments. Additionally, all steers were vaccinated against bovine rhinotracheitis virus, bovine viral diarrhea virus (types 1 and 2), bovine parainfluenza-3 virus, and bovine respiratory syncytial virus (Vista 5 SQ; Merck Animal Health, Kenilworth, NJ), clostridial pathogens (Vision 8 with Spur; Merck Animal Health), and Mycoplasma bovis (Myco-B ONE DOSE; American Animal Health, Inc. Grand Prairie, TX). Steers also received ivermectin (Vetrimec pour-on; Vet One, Boise, ID) for treatment of internal and external parasites. Steers were sorted by BW within arrival block to experimental treatment assignments. Treatments were then applied randomly to pens. On day 0, BW was collected, and this BW was averaged with the day −1 BW to calculate initial BW.

**Experimental Treatments**

Four treatments were used in a generalized complete block experimental design. The treatments consisted of the following: 1) a negative control, 5 mL of sterile saline injected subcutaneously (CON); 2) subcutaneous administration of florfenicol (Nuflor; Merck Animal Health) at 40 mg/kg of BW (NUF); 3) subcutaneous administration of ceftiofur in the posterior aspect of the ear (Excede; Zoetis, Parsippany, NJ) at 6.6 mg/kg of BW (EXC); and 4) subcutaneous administration of tulathromycin (Draxxin; Zoetis) at 2.5 mg/kg of BW (DRA). All antimicrobials were administered according to the label recommendations. A 3-day postmetaphylactic interval (PMI) was implemented for steers that received florfenicol, a 5-day PMI was implemented for steers that received ceftiofur, and a 7-day PMI was implemented for steers that received tulathromycin. Steers in the CON group did not have a PMI and were eligible for therapeutic treatment on day 0. Steers were observed daily for symptoms of BRD and assigned a clinical illness score (0–4 severity scale) as described by Pillen et al. (2016). The eligibility of steers for therapeutic treatment was determined by ear tag identification and treatment record by a treatment-blinded investigator. Steers were considered to have a clinical case of BRD if they met the following criteria: 1) animal had a clinical illness score of 2 and a rectal temperature of ≥37.5 °C or 2) a clinical illness

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**Table 1. Chronology of key events for high-risk steers in blocks 1 and 2.**

| Item                  | Block 1     | Block 2     |
|-----------------------|-------------|-------------|
| Receiving period      |             |             |
| Body weight measurement | Days -1, 0, 25, 38 | Days -1, 0, 33, 42 |
| Period 1              | Day 0 to 25 | Day 0 to 33 |
| Period 2              | Day 26 to 38 | Day 33 to 42 |
| Overall               | Day 0 to 38 | Day 0 to 42 |
| Finishing period      |             |             |
| Body weight measurement | Days 38, 252 | Days 42, 242 |
| Blood collection      |             |             |
| Initial               | Day 0       | Day 0       |
| Interim               | Day 126     | Day 123     |
| Final                 | Day 252     | Day 242     |

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1The length of the receiving period was not the same number of days for blocks 1 and 2. The cattle arrived on different days of the week and it was not possible for the receiving period to be an equal number of days between blocks 1 and 2.

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**Table 2. Diet formulation and composition of receiving and finishing diets fed to high-risk beef steers.**

| Item             | Receiving diet | Finishing diet |
|------------------|----------------|----------------|
| Ingredient, % DM |                |                |
| Steam-flaked corn| 20.63          | 65.06          |
| Sweet bran       | 53.46          | 20.39          |
| Chopped alfalfa  | 19.92          | 8.06           |
| Yellow grease    | 1.82           | 2.17           |
| Supplement       | 2.05           | 2.01           |
| Limestone        | 2.12           | 1.74           |
| Urea             | 0.57           |                |

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1Dry matter basis, except DM %.
2Supplement supplied 5.99% potassium chloride, 44.40% crude protein, 3.82% sodium, 8.34 mg/kg cobalt carbonate, 395.00 mg/kg copper sulfate, 408.00 mg/kg iron sulfate, 764 mg/kg manganese oxide, 2.92 mg/ kg selenium, 2,490.00 mg/kg zinc sulfate, and 30 g/t manganese sodium (Rumensin 96; Elanco Animal Health, Greenfield, IN) on a DM basis. Actual diet formulation based on weekly DM determinations.
3Analysis performed by Servi-Tech Laboratories, Amarillo, TX.
4Ne4, and Ne4 reported as tabular values (NASEM, 2016).
Metaphylactic antibiotics and growth of beef cattle

The pens had 15.95 m² of pen space and 60 cm of linear bunk space per animal. Previous metaphylactic treatments were tildipirosin at 4 mg/kg of BW (Zuprevo, Merck Animal Health) and were implanted with 36 mg of bovine viral diarrhea virus (types 1 and 2), bovine parainfluenza-3 virus, and bovine respiratory syncytial virus (Vista 5 SQ; Merck Animal Health) and were implanted with 36 mg of zeranol (Ralgro Merck Animal Health). If steers continued to be symptomatic after their third treatment, they were classified as chronic and removed from the study after the receiving period (N = 2, 1 steer from CON and 1 steer from NUF). Steers treated once for BRD were considered BRD1, steers treated twice for BRD were considered BRD2, and steers treated three-times for BRD were considered BRD3. Two steers (both from the EXC treatment group) were treated for BRD in the finishing period (after day 46), all others were treated in the receiving period (before day 46). In addition, four steers died during the finishing period (1 steer from DRA [bloat], 1 steer from EXC [1 bloat, 1 injury], and 1 from NUF [bloat]).

### Housing and Management

During the receiving period, steers were housed in partially shaded soil-surfaced pens with 57.6–63.4 m² of pen space and 44.3–48.8 cm² of linear bunk space per steer [Block 1 (N = 123 steers, 3 replications per treatment, 10–11 steers per pen); Block 2 (N = 115 steers, 3 replications per treatment, 10–11 steers per pen)]. All steers were fed the same receiving diet (Table 2), throughout the receiving period. Days to the first therapeutic treatment and rectal temperatures were summarized from individual treatment records and averaged by pen. The morbidity for BRD1, BRD2, BRD3, percent chronic, and respiratory mortality were summed by pen and divided by total number of steers in the pen during the receiving period. Days to the first therapeutic treatment and rectal temperatures were summarized from individual treatment records and averaged by pen. All data, with the exception of morbidity and mortality are reported with mortalities excluded.

### Calculations

The ADG and G:F were calculated on a live BW basis. Average daily gain was calculated by subtracting the initial BW from the final BW, then divided by days on feed. The G:F was computed as the quotient of ADG divided by daily DM. The carcass-adjusted data were calculated from the HCW divided by the overall average dressing percent.

### Statistical Analysis

Data were analyzed as a generalized complete block design using PROC MIXED in SAS (SAS Institute Inc., Cary, NC). Growth performance was analyzed separately in the receiving period and finishing period because the cattle were
Table 3. Effect of metaphylaxis with a sterile saline (negative control), tulathromycin, ceftiofur, and florfenicol on health outcomes of high-risk, newly received beef steers1

| Item                              | CON  | DRA  | EXC  | NUF  | SEM3 | P-value |
|-----------------------------------|------|------|------|------|------|---------|
| N, steers                         | 55   | 61   | 60   | 62   | -    | -       |
| N, pens                           | 6    | 6    | 6    | 6    | -    | -       |
| BRD11 %                           | 58.8 | 26.3 | 26.3 | 45.2 |       | <0.01   |
| BRD22 %                           | 29.3 | 3.3  | 8.4  | 14.5 |       | <0.01   |
| BRD3%                             | 7.4  | 0    | 1.7  | 1.7  | 0.20  |         |
| Chronic6 %                        | 1.9  | 0    | 0.1  | 1.6  | 0.57  |         |
| Respiratory mortality %           | 1.8  | 1.6  | 1.7  | 0    | 1.55  | 0.80    |

1 a,b,Within a row, means without a common superscript differ, P < 0.05.
2 CON = no metaphylaxis at arrival; DRA = tulathromycin (Draxxin, Zoetis) administered as metaphylaxis at arrival; EXC = ceftiofur (Excede, Zoetis) administered as metaphylaxis at arrival; NUF = florfenicol (Nuflor, Merck) administered as metaphylaxis at arrival.
3 Poole standard error of least squares mean (N = 6 pens/mean).
4 Percentage of steers treated for bovine respiratory disease (BRD) at least once.
5 Percentage of steers treated for BRD at least twice.
6 Percentage of steers treated for BRD at least three times.
7 Percentage of steers treated for BRD four or more times.

re-penned after the completion of the receiving period. For growth performance during the receiving and finishing period, pen was the experimental unit, and the model included the fixed effect of metaphylactic treatment and the random effect of arrival block. Morbidity and mortality were analyzed as a binomial proportion using the GLIMMIX procedure of SAS. The CBC data were analyzed with repeated measures using animal as the experimental unit. The model included metaphylactic treatment × day, and the interaction of metaphylactic treatment × day. Animal within pen was the subject of the repeated measures and was included to control for any pen variation present when re-allotting cattle between the growing and finishing phase. Multiple covariance structures were tested, and the autoregressive 1 covariance structure resulted in the smallest Akaake and Schwarz Bayesian criteria and was considered the most appropriate for repeated measure analyses. Main effect means are presented for most CBC variables and in the presence of an interaction (P < 0.05) simple effect means are discussed in the text. For all variables, statistical significance was determined at P ≤ 0.05 and tendencies were noted at 0.05 < P ≤ 0.10.

RESULTS AND DISCUSSION

Morbidity and Mortality

Health outcomes during the receiving period are presented in Table 3. A greater (P < 0.01) percentage of steers were treated for BRD1 in the CON and NUF treatments than DRA and EXC. The lack of difference in BRD1 treatments among CON and NUF steers is similar to observations by Martín et al. (2007). Nonetheless, these results contrast with Catry et al. (2008), where metaphylactic florfenicol (NUF) produced fewer subsequent treatments for BRD than cattle that did not receive metaphylaxis. In addition, Gonzalez-Martin et al. (2011) reported no difference in BRD incidence among cattle receiving florfenicol (NUF) or tulathromycin (DRA) as metaphylaxis. The disparity between the current experiment and those of Gonzalez-Martin et al. (2011) and Catry et al. (2008) may be location, season, or type of pens, or the initial health status of the cattle. Similarly, Catry et al. (2008) and Gonzalez-Martin et al. (2011) used dairy calves and excluded any calves that appeared morbid during the experimental animal selection process. In the present study, no steers were excluded from the experiment based on arrival health status. Likewise, both studies began when a defined BRD outbreak occurred, rather than assigning metaphylactic treatment after feedlot arrival as in the present study.

In the present experiment, DRA decreased BRD1 by 42% compared to NUF (P = 0.04) and 55% compared to CON (P < 0.01), which is consistent with previous studies using high-risk cattle (Godinho et al., 2005; Kilgore et al., 2005; Nutsch et al., 2005; Rooney et al., 2005; Skogerbø et al., 2005; Tennant et al., 2014). Moreover, EXC decreased BRD2 by 55% compared to CON (P < 0.01), which aligns with previous studies (Hibbard et al. 2002a, 2002b; Encinas et al., 2006). As expected, a greater percentage of CON steers were treated for BRD2 than steers given antimicrobial metaphylaxis (P < 0.01), similar to reports by Godinho et al. (2005) where a greater proportion of cattle receiving no metaphylaxis required additional therapeutic treatment for BRD. The DRA treatment tended (P = 0.08) to decrease morbidity rate of BRD2 by 77% compared to NUF, and EXC did not differ (P ≥ 0.27) from NUF and DRA treatment groups. There was a tendency (P = 0.10) for CON steers to have a greater percentage of BRD3 treatments. Nevertheless, the percentage of chronic steers and respiratory mortality did not differ (P > 0.57).

The EXC group had the greatest days to first therapeutic treatment, whereas CON had the fewest (P < 0.01), with DRA and NUF being intermediate. There was no difference in days to second or third therapeutic treatment (P ≥ 0.26). These results could be confounded with the difference in PMI among metaphylactic antimicrobials used and design of the experiment thus results should be interpreted with caution. The DRA treatment had a 7-d PMI, the EXC treatment had a 5-d PMI, the NUF treatment had a 3-d PMI, and steers in the CON treatment could be therapeutically treated on day 0. Indeed, tulathromycin is known to be slowly metabolized in the body and have a prolonged activity in the lungs, and therefore it would be expected to have the greatest days to first therapeutic treatment (Evans, 2005), which was not the case in the present study. Cefotiofur is reported to have a half-life of 6 h, which is appreciably longer than most other cephalosporins in part because of the administration route in the postauricular aspect of the ear extending the duration of action (Merck, 2021). However, this does not explain the difference in days to first treat between the EXC and DRA treatment groups.

Rectal temperature did not differ (P > 0.53) among groups during any therapeutic treatment which is consistent with previously published literature (Godinho et al., 2005; Munoz et al., 2020). Nonetheless, Catry et al. (2008) reported rectal
temperature was greater for calves given florfenicol than calves given no metaphylaxis. Rectal temperature is commonly used as a proxy for core body temperature to objectively diagnose illness and disease and is reportedly accurate in clinical diagnosis of BRD (Gonzalez-Martin et al., 2011).

Receiving Growth Performance

Growth performance during the receiving period is reported in Table 4. There were no differences in initial (P = 0.99) or final BW among treatments (P = 0.63). The literature surrounding the effect of metaphylaxis on BW within the receiving period is inconsistent. Our findings are supported by studies comparing tulathromycin and cephalothin as metaphylaxis to a negative control that reported no difference in final BW (Godinho et al., 2005; Benton et al., 2008). In contrast, others have reported increased final BW during the receiving period when metaphylaxis is given (Catry et al., 2008; Munoz et al., 2020). The inconsistency of results could be explained by a different length of time in the receiving period [20 d in the Catry et al. (2008) study, and 56 d in the Munoz et al. (2020) study, compared to 38–45 d in the present study].

In period 1, steers administered DRA had greater ADG than other treatments (P < 0.01). Steers given DRA had a 49% increase (P < 0.01) in overall ADG compared to the negative control. Purportedly, cattle experiencing active BRD have less ADG, thus because the DRA morbidity in the current experiment was least among treatments, it is plausible that this is the reason the ADG was greatest among treatments (Griffin, 1997; Gardner et al., 1999; Munoz et al., 2020). The NUF treatment had the least ADG within period 1 (P < 0.01). These data agree with Boyd et al. (2006), where lightweight calves were given a negative control, cephalothin, or florfenicol as metaphylaxis on day 0. The ADG of calves treated with florfenicol was less than control and cephalothin groups, which corresponded to an increase in BRD1 for steers given florfenicol vs. control or cephalothin at arrival and is likely the reason ADG was less. In the present experiment, NUF had greater ADG in period 2 (P = 0.05) than CON and DRA but did not differ from EXC (P = 0.35). This is likely because of compensatory gain in response to a decreased ADG in period 1. The overall ADG for the receiving period was greatest for DRA, and least for the steers treated metaphylactically with NUF (P < 0.01), with CON and EXC being intermediate.

Period 1 DMI was greatest in DRA and EXC, and least in CON and NUF (P < 0.01). Florfenicol has been reported to temporarily reduce DMI of cattle (Boyd et al., 2006; Catry et al., 2008). In the current study, DRA tended (P = 0.08) to have a greater DMI than NUF in period 2. As expected, based on the results of periods 1 and 2 combined, the overall DMI was greatest for DRA and least for NUF (P = 0.02). Our findings did not indicate that overall DMI during the receiving period for the CON treatment differed from steers that received metaphylaxis at arrival. In studies by Word et al. (2021) and Tennant et al. (2014), cattle receiving metaphylaxis did not have a greater DMI than the cattle administered a negative control. Conversely, others reported metaphylaxis increased DMI during the feedlot receiving period (Munoz et al., 2020).

There was a tendency (P = 0.08) for DMI as a percent of BW to differ in period 1. In addition, there was no difference in DMI as a percent of BW in period 2 or for the overall (P ≥ 0.14) receiving period, which suggests that even though a tendency for a difference was present early after feedlot arrival (period 1), the steers with decreased DMI compensated by the end of the receiving period.

The G:F in period 1 was greatest for DRA, least for NUF, with CON and EXC being intermediate and not differing (P < 0.01). Tulathromycin resulted in a greater G:F than CON, which was likewise observed by Munoz et al. (2020). Further, our finding that florfenicol resulted in the least G:F has been observed previously (Boyd et al., 2006). Others have reported that cephalothin increased G:F over a negative control (Benton et al., 2008). In period 2, the G:F was greatest (P = 0.01) for the steers given NUF; perhaps because of compensatory gain after a decreased DMI and G:F in period 1. Likewise, EXC tended (P = 0.06) to differ from NUF; whereas CON and DRA had less G:F than NUF (P < 0.01) and did not differ in G:F during period 2. For the overall receiving period, G:F was greatest (P = 0.03) in steers that received DRA; whereby, CON, EXC and NUF did not differ (P ≥ 0.07). The increased G:F observed in the DRA treatment group in the present study is in contrast with the receiving period data reported by Gonzalez-Martin et al. (2011), though when G:F was evaluated over the entire receiving and finishing period no difference in G:F was reported.
Finishing Period Growth Performance and Carcass Characteristics

Growth performance during the finishing period is presented in Table 5. There was no difference among treatments in the initial BW ($P = 0.33$) or final BW ($P = 0.22$). Metaphylaxis at arrival generally does not change final BW before harvest (Gardner et al., 1999; Skogerboe et al., 2005; Booker et al., 2006; Word et al., 2021). Conversely, a study by Tennant et al. (2014) reported that final BW was greater in steers given tulathromycin as metaphylaxis at arrival compared to steers not given metaphylaxis. Although not statistically different, DRA and EXC both numerically increased final BW over CON and NUF; whereby an increase of 20 kg or greater in final BW is biologically relevant.

Over the entire finishing period, ADG was greatest for EXC, and least for CON ($P = 0.05$). In addition, the present study reported a 3.4% greater ADG between the DRA and CON treatment groups which is less than Munoz et al. (2020) reported, where metaphylactic tulathromycin resulted in a 15.8% greater ADG than a negative control. Previous literature on whether ADG during the finishing period is affected by arrival metaphylaxis is inconsistent. It has been commonly reported that metaphylaxis at arrival has no effect on ADG (Griffin, 1997; Gonzalez-Martín et al., 2011; Word et al., 2021). Additionally, a study by Tennant et al. (2014) reported cattle given tulathromycin as metaphylaxis had greater ADG than cattle given a negative control, which is in contrast to the present study where the CON and DRA treatments did not differ ($P = 0.19$) in ADG during the finishing period. The difference between these results could be explained by the overall decreased morbidity across the entire experimental period in both the steers given tulathromycin and the steers given a negative control in the Tennant et al. (2014) study. Furthermore, others have reported a greater ADG in cattle given tulathromycin as metaphylaxis than those given florfenicol (Rooney et al., 2005; Skogerboe et al., 2005), which is in contrast to the present study where DRA and NUF did not differ ($P = 0.70$) in overall ADG during the finishing period. Nonetheless, a study conducted by Gardner et al. (1999) reported cattle that exhibited clinical signs of BRD or those that had lung lesions at harvest had a decreased ADG than healthy cattle, which agrees with our finding that the EXC treatment had a greater ADG than the CON treatment.

There was no difference ($P = 0.14$) in DMI during the overall finishing period, which agrees with previous literature (Tennant et al., 2014; Word et al., 2021). Likewise, DMI as a percent of BW did not differ ($P = 0.23$) among treatment groups. There is some evidence that administering ceftiofur as metaphylaxis increases DMI in the finishing period. In a study by Booker et al. (2006), calves given ceftiofur as metaphylaxis had a greater DMI over the entire finishing period than those given tilmicosin phosphate (Booker et al., 2006). The G:F among all treatments in the finishing period did not differ ($P = 0.40$). Our findings that metaphylaxis at arrival had no impact on G:F in the finishing period have been previously noted (Rooney et al., 2005; Booker et al., 2006; Gonzalez-Martín et al., 2011; Tennant et al., 2014; Word et al., 2021).

Carcass-adjusted ADG followed the same trend as ADG calculated from live BW where ADG was greatest for EXC and least for CON ($P = 0.05$; Table 5). No differences in carcass-adjusted growth performance, carcass characteristics, or liver score were observed (Table 6; $P \geq 0.12$). These results are in agreement with prior terminal works that evaluated differing metaphylactic drugs and demonstrate that cattle treated for BRD multiple times are able to achieve similar compositional endpoints as nontreated contemporaries (Wilson et al., 2017; Word et al., 2021).

Complete Blood Count

The least squares means for initial, interim, and final CBC variables by day are presented in Table 7. All variables were affected by day ($P < 0.01$) except for mean corpuscular hemoglobin concentration ($P = 0.29$).

There was a treatment × time interaction for platelet count ($P = 0.02$; Figure 1). Steers administered CON had greater concentrations of platelets than DRA or EXC at the initial measurement ($P < 0.01$), though at this time metaphylactic treatment had yet to be administered and the reason for the difference in baseline is unknown. Platelets decreased in the steers administered DRA as metaphylaxis between interim and final ($P < 0.01$). Platelets are responsible for blood clotting and high platelet count can indicate inflammation, which is plausible as platelet count was greatest for all treatments at the initial time point, when stress and inflammation were likely increased as a result of the stress of marketing and transportation. To mediate inflammation, platelets exert chemotaxis on leukocytes (Klinger, 1997). In the present study,

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**Table 5.** Effect of metaphylaxis with a sterile saline (negative control), tulathromycin, ceftiofur, and florfenicol on finishing growth performance of high-risk, steers

| Item                        | Treatment† | P-value |
|-----------------------------|------------|---------|
| N, steers                   | CON 55     |        |
| N, pens                     | DRA 61     |        |
|                             | EXC 60     |        |
|                             | NUF 62     |        |
|                             | SEM        | -       |
|                             | P-value    | -       |
| Initial body weight†, kg    | CON 294    |        |
|                             | DRA 306    |        |
|                             | EXC 299    |        |
|                             | NUF 288    |        |
|                             | 7.03       | 0.33    |
| Final body weight†, kg      | CON 581    |        |
|                             | DRA 602    |        |
|                             | EXC 607    |        |
|                             | NUF 587    |        |
|                             | 14.9       | 0.22    |
| Overall finishing           |            |         |
| ADG, kg                     | CON 1.41a  |         |
|                             | DRA 1.46b  |         |
|                             | EXC 1.52c  |         |
|                             | NUF 1.48d  |         |
|                             | 0.030      | 0.05    |
| DMI, kg                     | CON 8.47   |         |
|                             | DRA 8.59   |         |
|                             | EXC 8.97   |         |
|                             | NUF 8.47   |         |
|                             | 0.172      | 0.14    |
| DMI, % of BW                | CON 1.94   |         |
|                             | DRA 1.89   |         |
|                             | EXC 1.99   |         |
|                             | NUF 1.94   |         |
|                             | 0.031      | 0.23    |
| G:F                         | CON 0.167  |         |
|                             | DRA 0.171  |         |
|                             | EXC 0.170  |         |
|                             | NUF 0.175  |         |
|                             | 0.0032     | 0.40    |
| Carcass-adjusted            |            |         |
| Final BW†, kg               | CON 586    |         |
|                             | DRA 606    |         |
|                             | EXC 606    |         |
|                             | NUF 590    |         |
|                             | 13.6       | 0.20    |
| ADG, kg                     | CON 1.45a  |         |
|                             | DRA 1.48b  |         |
|                             | EXC 1.52c  |         |
|                             | NUF 1.50d  |         |
|                             | 0.027      | 0.05    |
| DMI, % of BW                | CON 1.93   |         |
|                             | DRA 1.88   |         |
|                             | EXC 1.98   |         |
|                             | NUF 1.93   |         |
|                             | 0.028      | 0.24    |
| G:F                         | CON 0.171  |         |
|                             | DRA 0.172  |         |
|                             | EXC 0.169  |         |
|                             | NUF 0.175  |         |
|                             | 0.031      | 0.40    |

†Within a row, means with different superscripts differ $P < 0.05$.

1CON = no metaphylaxis at arrival; DRA = tulathromycin (Draxxin, Zoetis) administered as metaphylaxis at arrival; EXC = ceftiofur (Excide, Zoetis) administered as metaphylaxis at arrival; NUF = florfenicol (Nuflox, Merck) administered as metaphylaxis at arrival.

2Standard error of least squares mean ($N = 6$ pens per mean).

3All BW are shrunk by 4%.

4Hot carcass weight divided by overall average DP (65.8%).

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Table 6. Carcass characteristics and liver score of finishing beef steers administered metaphylaxis with tulathromycin, ceftiofur, florfenicol, or given no metaphylaxis at feedlot arrival

| Item                        | CON   | DRA   | EXC   | NUF   | SEM  | P-value |
|-----------------------------|-------|-------|-------|-------|------|---------|
| Hot carcass weight, kg      | 385   | 399   | 398   | 388   | 10.7 | 0.44    |
| DP, %                       | 65.78 | 65.75 | 65.69 | 65.73 | 0.434| 0.99    |
| Marbling score              | 553   | 520   | 532   | 545   | 18.1 | 0.45    |
| Fat thickness, cm           | 1.74  | 1.77  | 1.74  | 1.75  | 0.105| 0.99    |
| Longissimus dorsi area, cm^2| 90.41 | 91.64 | 92.83 | 90.25 | 2.454| 0.66    |
| Calculated YG               | 3.46  | 3.54  | 3.45  | 3.49  | 0.166| 0.98    |
| EBP, %                      | 31.82 | 31.85 | 31.75 | 31.85 | 0.670| 0.99    |
| AFBW, kg                    | 549   | 568   | 570   | 551   | 14.8 | 0.18    |
| Choice or greater, %        | 82.4  | 87.8  | 86.5  | 92.2  | 4.70 | 0.51    |
| Abscessed livers, %         | 17.7  | 12.7  | 15.6  | 14.1  | 9.76 | 0.93    |

Table 7. Day effects of complete blood count for high-risk beef steers given metaphylaxis with a negative control, tulathromycin, ceftiofur, and florfenicol at arrival

| Item                              | Initial | Interim | Final | SEM  | P-value |
|-----------------------------------|---------|---------|-------|------|---------|
| Red blood cells, M/µL             | 10.13a  | 9.97b  | 9.13c | 0.067| <0.01   |
| Hemoglobin, g/dL                  | 13.24a  | 14.64b | 14.91c| 0.083| <0.01   |
| Hematocrit, %                     | 39.74a  | 43.97b | 44.67c| 0.271| <0.01   |
| Mean corpuscular volume, fl       | 39.49a  | 44.15b | 49.14c| 0.286| <0.01   |
| Mean corpuscular hemoglobin, pg   | 13.19a  | 14.70b | 16.40c| 0.093| <0.01   |
| Mean corpuscular hemoglobin%      | 33.43   | 33.21  | 33.40 | 0.127| 0.29    |
| Reticulocyte, K/µL                | 0.62c   | 1.78c  | 1.82c | 0.146| <0.01   |
| Reticulocyte, %                   | <0.01c  | 0.01c  | 0.01c | 0.003| 0.01    |

WBC were elevated and could explain why platelets were greatest initially in correlation with the greater morbidity rate, though this does not offer an explanation as to why the CON treatment had greater platelet count than DRA or EXC. Furthermore, excessive platelet count is uncommon and likely a secondary condition observed with stress (Roland et al., 2014), which steers were experiencing at the initial measurement. The literature about the significance of platelets as an immune response to BRD is inconsistent. Platelets have often been reported as poor predictors of BRD (Richeson et al., 2013; Moisá et al., 2019). In a study by Fontenot (2015), platelets were reported to decrease with the number of times calves were treated for BRD.

An elevated red blood cell (RBC) count can indicate lung disease, inflammation, or dehydration; conditions which are routinely associated with high-risk calves at feedlot arrival. Likewise, elevated RBC count can indicate a greater risk of BRD diagnosis (Richeson et al., 2013). In the present study, RBC count was decreased at the final compared to initial and interim measurements (P < 0.01); however, all values observed during the study were within normal ranges for bovine (Merck, 2021). Hemoglobin concentration, hematocrit, mean corpuscular volume, and mean corpuscular hemoglobin increased (P < 0.01) across all time points, where initial was least, interim was intermediate, and final was greatest with no abnormal values observed (Merck, 2021). Likewise, reticulocyte count and percentage increased from initial to interim (P ≤ 0.01). These RBC-related measurements are all closely related and the pathogenic state causing an abnormal value of one variable is often related to the same of another. For example, decreased hemoglobin at feedlot entry has been correlated to decreased incidence of BRD treatment (Fontenot, 2015). Closely related to RBC and hemoglobin, hematocrit is the ratio of RBC, leukocytes, and platelets to total blood volume (Pagana et al., 2015). Hematocrit is often used as a proxy to indicate dehydration. Overall, it is important to note that hematological ranges fluctuate with age and the time effects in which differentials change but remain within a normal range could be confounded with age and the length of time in between sample collections (Roland et al., 2014).

The least squares means for metaphylactic treatment effects on CBC variables are presented in Table 8. Metaphylactic treatment had no effect (P ≥ 0.17) on any blood variables throughout the study. This is likely confounded by sample time, as treatment had not been administered when the first sample was collected, and steers morbidity was nonexistent at the other two sample points. If blood had been collected at times when steers were experiencing the greatest morbidity...
The neutrophil:lymphocyte ratio decreased ($P < 0.01$) from initial to interim and increased ($P < 0.01$) from interim to final, though it was within normal ranges. Lymphocytes are the most abundant subpopulation of WBC in bovines, therefore a ratio greater than 1 is an indicator of stress (Roland et al., 2014). Similar to RBC and their differentials, changes in WBC and their differentials that do not change outside of normal ranges are likely because of age and are confounded with sampling times. The metaphylactic treatment effects on WBC differentials are presented in Table 10. There was a tendency ($P = 0.10$) for EXC to have a greater neutrophil count than NUF, though EXC did not differ ($P \geq 0.06$) from DRA or CON. There was no treatment effect ($P \geq 0.12$) on lymphocyte concentration, eosinophil concentration and percentage, neutrophil percentage, basophil percentage, and neutrophil:lymphocyte ratio. Additionally, these differentials were all within normal ranges (Merck, 2021).

Lindholm-Perry et al. (2018) reported that neutrophils were greater in morbid cattle compared to those that were healthy or asymptomatic. It is unclear why the EXC treatment would have greater concentrations of neutrophils than the NUF treatment when the morbidity rate in the NUF treatment group was greater. It is important to note that neutrophils have been reported to be poor predictors of BRD (Richeson et al., 2013), suggesting the treatment differences are better explained by the BRD incidence rates of the antibiotics used as metaphylaxis than as a result of immune response to BRD.

A treatment × time interaction was observed for WBC counts ($P < 0.01$; Figure 2). The CON treatment had the least WBC at the interim measurement compared to DRA, EXC, and NUF ($P \leq 0.01$). Steers in the EXC and NUF treatment group had a decrease in WBC across all time points ($P \leq 0.01$) compared to CON and DRA treatments. A greater WBC concentration can be indicative of infection, inflammation, or stress (Pagana et al., 2015). The greatest WBC count on day 0 is expected as calves had recently undergone transport and handling stress as well as a greater morbidity rate than the other two 2 time points.

There was a treatment × time interaction for monocyte concentrations ($P < 0.01$; Figure 3). At the interim measurement, steers administered CON, DRA, and NUF had lesser after metaphylaxis treatment was administered, the results could be very different.

The least squares means for initial, interim, and final day effects on WBC differentials are presented in Table 9. All variables were affected by day ($P < 0.01$), but nonetheless all observed values were within normal reference ranges. Neutrophil count was greater at the initial time point and decreased at the interim and final measurements. According to Pagana et al. (2015), a greater neutrophil count can indicate inflammation, infection, or stress, and they are particularly implicated in the pathogenesis of inflammatory respiratory tract diseases (Earley et al., 2016), which could explain why the observed neutrophil count was greater initially among treatments. The lymphocyte and eosinophil concentrations decreased from interim to final ($P < 0.01$) while basophil count decreased from initial to interim ($P < 0.01$) and eosinophil percentage increased ($P < 0.01$) from initial to interim. The neutrophil:lymphocyte ratio decreased ($P < 0.01$) from initial to interim and increased ($P < 0.01$) from interim to final, though it was within normal ranges. Lymphocytes are the most abundant subpopulation of WBC in bovines, therefore a ratio greater than 1 is an indicator of stress (Roland et al., 2014). Similar to RBC and their differentials, changes in WBC and their differentials that do not change outside of normal ranges are likely because of age and are confounded with sampling times. The metaphylactic treatment effects on WBC differentials are presented in Table 10. There was a tendency ($P = 0.10$) for EXC to have a greater neutrophil count than NUF, though EXC did not differ ($P \geq 0.06$) from DRA or CON. There was no treatment effect ($P \geq 0.12$) on lymphocyte concentration, eosinophil concentration and percentage, neutrophil percentage, basophil percentage, and neutrophil:lymphocyte ratio. Additionally, these differentials were all within normal ranges (Merck, 2021).

Lindholm-Perry et al. (2018) reported that neutrophils were greater in morbid cattle compared to those that were healthy or asymptomatic. It is unclear why the EXC treatment would have greater concentrations of neutrophils than the NUF treatment when the morbidity rate in the NUF treatment group was greater. It is important to note that neutrophils have been reported to be poor predictors of BRD (Richeson et al., 2013), suggesting the treatment differences are better explained by the BRD incidence rates of the antibiotics used as metaphylaxis than as a result of immune response to BRD.
Monocyte concentrations than EXC ($P \leq 0.01$). Likely, NUF and CON had a greater monocyte percentage as a result of the greater morbidity rates compared to the other treatments. Like neutrophils, monocytes are phagocytic, and a greater monocyte count or percentage can be indicative of inflammation or viral infection (Roland et al., 2014; Pagana et al., 2015). Conversely, Lindholm-Perry et al. (2018) reported calves with a low monocyte count at weaning were more likely to develop BRD.

There was a treatment × time interaction for percentage of lymphocytes ($P = 0.03$; Figure 4). The CON group had a greater interim lymphocyte percentage than both DRA and EXC ($P \leq 0.04$). Lymphocytes are the largest proportion of WBC in cattle, though they decrease with age (Roland et al., 2014). In a study conducted by Moisá et al. (2019), a negative association was reported between lymphocyte concentration and BRD. Nevertheless, it should be noted that in other literature lymphocytes have been reported to be poor predictors of BRD based on a low area under the curve (Richeson et al., 2013).

**CONCLUSION**

The results of this study indicate administering metaphylaxis after arrival with either tulathromycin or ceftiofur can decrease the morbidity rate of high-risk calves. Administering florfenicol as metaphylaxis after arrival increased cattle treated for BRD once by 23.1% vs. a negative control where steers did not receive metaphylaxis. During the receiving period, the ADG of steers administered tulathromycin was greatest and DMI and G:F was greatest among those administered tulathromycin and ceftiofur. Tulathromycin increased ADG during the growing phase and ceftiofur had the greatest long-term increase in ADG vs. CON, but did not differ from other metaphylactic antimicrobials used. Bovine respiratory disease remains a complex and multifactorial health challenge in high-risk cattle, but producers can improve the health outcomes of their cattle with several commercially available antimicrobials administered as metaphylaxis that decrease morbidity and mortality and subsequently improve growth performance.

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**Conflict of interest statement**

None declared.
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