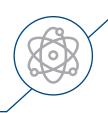
Research Notes ARM & HAMMER



More is not always better: Bacillus dose impact on immune gene expression response

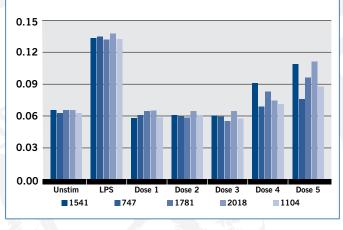
STUDY OVERVIEW

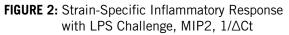
- A lab trial¹ was conducted to assess the effects of increasing doses of Bacillus strains on the inflammatory response of intestinal epithelial cells
- Proprietary ARM & HAMMER[™] Bacillus strains were titrated in one log increments to derive five dose levels and administered to IEC6 intestinal epithelial cell culture
- Strains were tested both with and without exposure to lipopolysaccharide (LPS) as an inflammatory challenge
- Gene expression of the inflammatory cytokine, macrophage inflammatory protein-2 (MIP2), was used as the indicator of inducing an inflammatory response

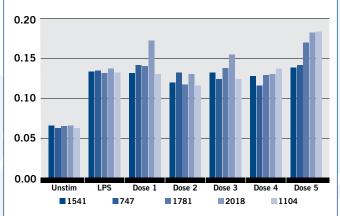
RESULTS

- Inflammatory responses were not induced by any Bacillus strains when administered at dose levels 1, 2 and 3 in cell culture
- Bacillus strains 1541 and 1781 increased inflammatory cytokine gene expression when administered to cell culture at dose levels 4 and 5 (Fig. 1)
- Bacillus strains 2018 and 1104 increased inflammatory cytokine gene expression when administered to cell culture at dose level 5 (Fig. 1)
- When combined with LPS, Bacillus strains 1781, 2018 and 1104 exacerbated the inflammatory response compared to LPS alone (Fig. 2)

FIGURE 1: Strain-Specific Inflammatory Response, MIP2, $1/\Delta Ct$







CONCLUSIONS

- Increasing doses of *Bacillus* generated inflammatory response in intestinal epithelial cells
- In application, greater doses of *Bacillus* do not always result in better responses; rather, the highest dose levels increased inflammatory response in this trial



1 Davis E, Christianson J. Immune Gene Expression Response to Dose Titrations of ARM & HAMMER Proprietary Bacillus Probiotics (CERTILLUS™). 2017. Report on file.



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