

Parabacteroides distasonis Alleviates Intestinal Barrier Injury Induced by ETEC

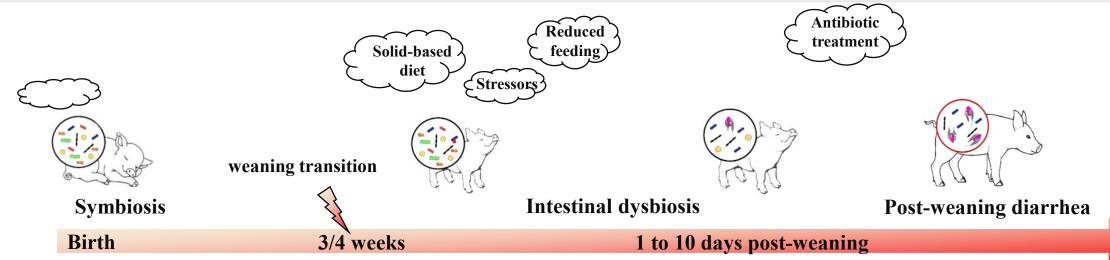
EAAP 2024 Florence Italy

Jing Wang, Bie Tan

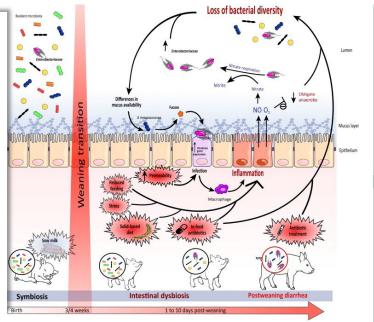
Sep 3th 2024



Gut microbiota dysbiosis → Post-weaning diarrhea



- **❖** Disrupt gut microbiota composition
- More susceptible to pathogenic
 bacteria, e.g. Enterotoxigenic
 Escherichia Coli (ETEC) → growth
 inhibition → diarrhea → death



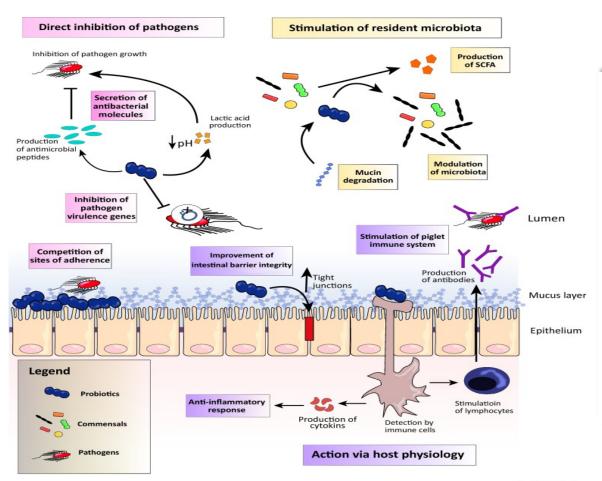


Gresse R, et al., Trends Microbiol, 2017

朴诚奋勉 求实创新



Probiotics restore gut microbial balance in weaning piglets



- ***** Balance the gut microbial homeostasis.
- ***** Competitive exclusion of pathogens and producing antimicrobial substances.
- **Stimulation of the host immune system** and improvement in intestinal barrier function.

Trends in Microbiolog

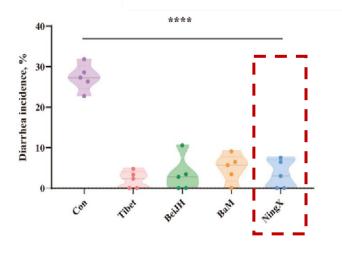


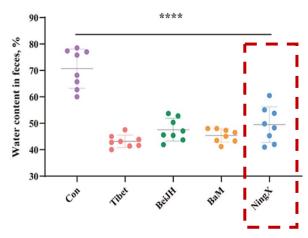
Ningxiang pig showed stronger diarrhea resistance

Ningxiang pig Native Chinese breeds

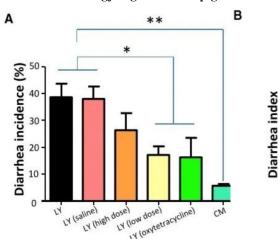


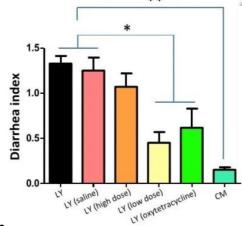
***** Lower diarrhea incidence





CM: Congjiang miniature piglets LY: Landrance × Yourshire piglets





 Microbiota from local piglet could target the LY piglet to confer diarrhea resistance

Hu J, et al., Cell Host & Microbe, 2018; Ma N, et al., Frontiers in Nutrition. 2022.

- I. What are the differential gut microbes in NingXiang piglets?
- II. Does the differential microbe regulate diarrhea resistance and gut barrier function?



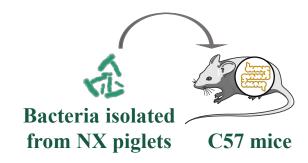
Trial I. Microbial screening

Ningxiang piglet Yorkshire piglet



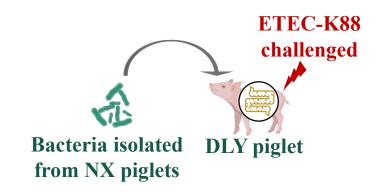
- Screen differential gut microbes
- Isolate the differential microbe

Trial II. Microbial function verification in mice



- Transplant the differential bacteria to mice
- Detect the intestinal development and barrier function

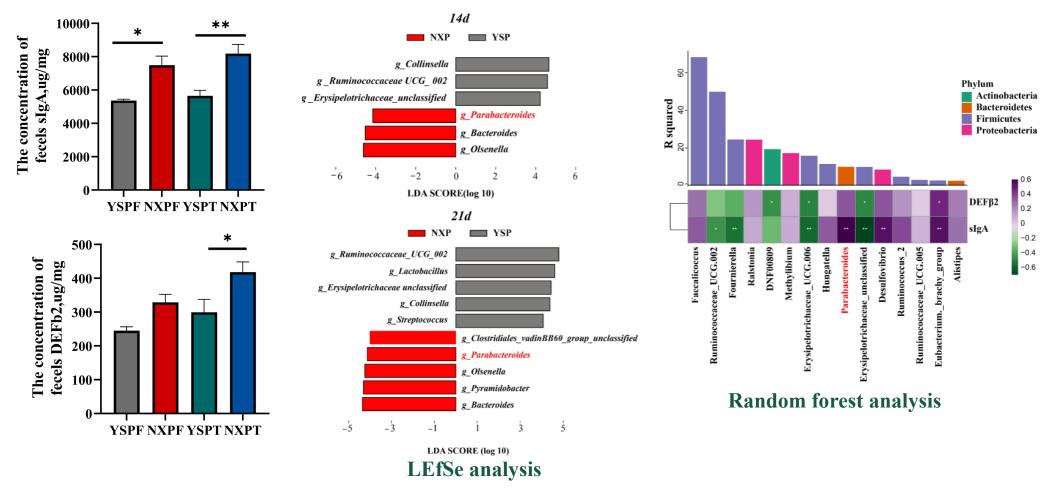
Trial II. Microbial function verification in DLY piglets



- Transplant the differential bacteria to DLY piglets
- Observe the diarrhea rate and intestinal injury induced by ETEC
- Analyze the gut microbial structure and composition



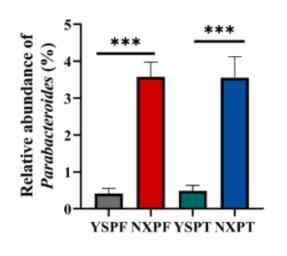
Parabacteroides were positively correlated with fecal sIgA

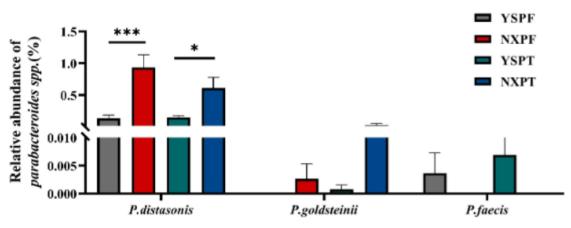


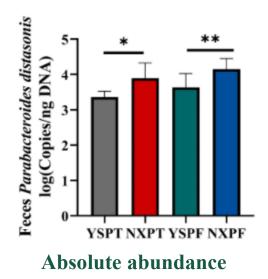
- ✓ The fecal sIgA and concentrations in Ningxiang piglets were higher than those of YS piglets.
- ✓ Parabacteroides were dominant in Ningxiang piglets and positively correlated with fecal sIgA.

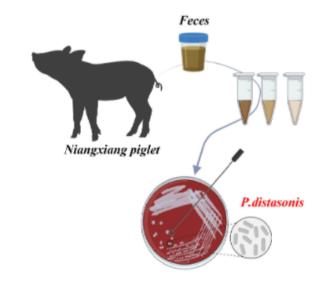


P. distasonis was isolated from the Ningxiang piglet's feces









- ✓ The abundance of *P. distasonis* was higher than *P. goldsteinii* and *P. faecis* in Ningxiang piglets.
- ✓ A higher presence of *P. distasonis* in Ningxiang piglets at 14 d and 21 d of age.



Trial I. Differential microbial screening

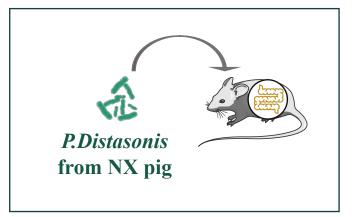
Ningxiang piglet Yorkshire piglet



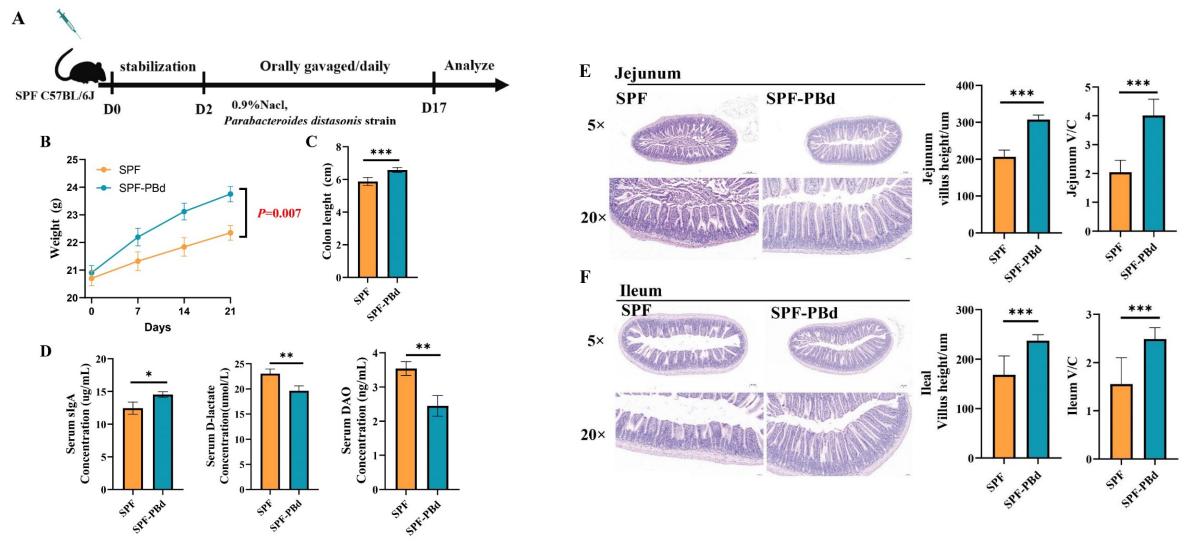
- ✓ Parabacteroides were dominant in Ningxiang piglets
- ✓ Parabacteroides were positively correlated with fecal sIgA
- ✓ *P. distasonis* was isolated from Ningxiang piglet feces.

Does *P. distasonis* have a beneficial effect on intestinal development or barrier function?



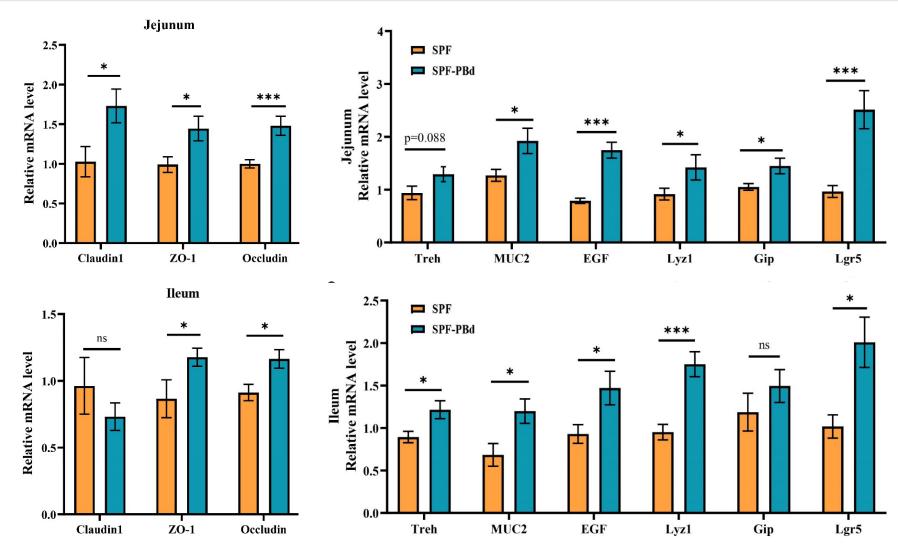


P. distasonis improved intestinal barrier integrity and development



- ✓ *P.distasonis* administration promoted the body weight gain of mice.
- ✓ *P.distasonis* increased the serum sIgA level and intestinal V/C ratio, and decreased the gut barrier permeability.

P. distasonis improved intestinal barrier integrity and development



- ✓ *P.distasonis* administration increased the intestinal tight junction gene expressions in mice.
- ✓ *P.distasonis* might promote the intestinal development.



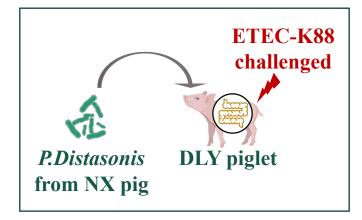
Trial II. Microbial function verification in mice



- ✓ *P. distasonis* increases the body weight of mice
- ✓ *P.distasonis* enhances the serum sIgA concentrations
- ✓ *P.distasonis* promotes the intestinal integrity and development

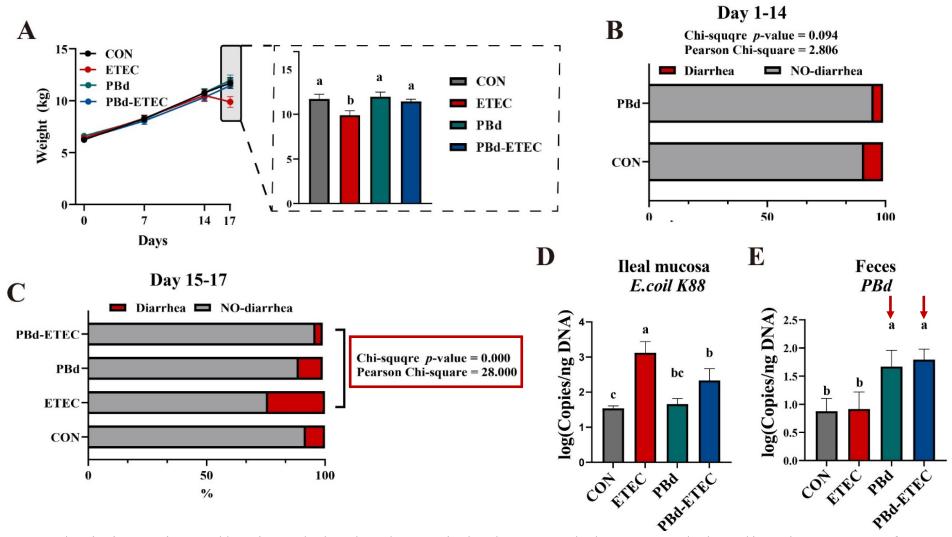
Does *P. distasonis* prevent diarrhea in DLY piglets?







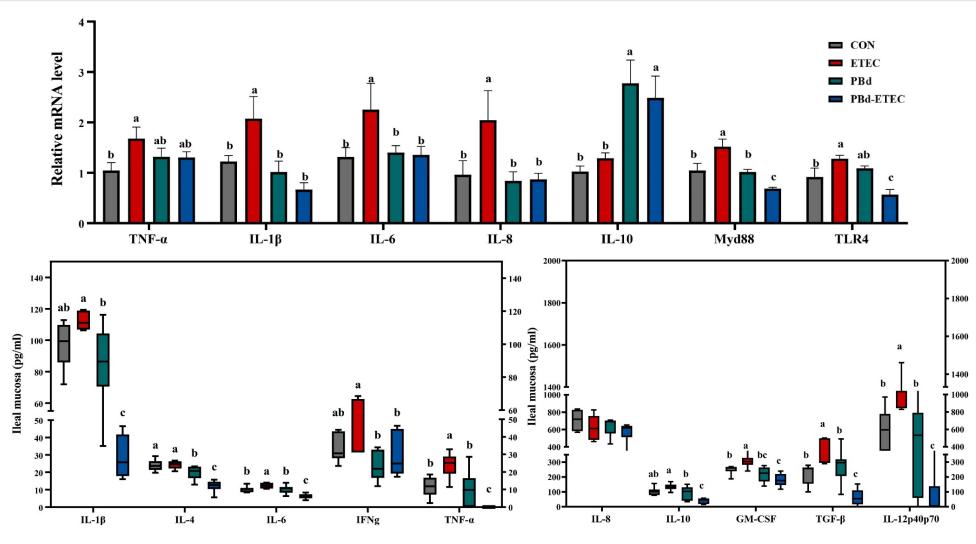
P. distasonis ameliorated diarrhea induced by ETEC infection



- ✓ *P.distasonis* administration alleviated the body weight loss and decreased the diarrhea rate after ETEC infection.
- ✓ *P.distasonis* could colonized in the gut of DLY piglets and inhibited the ETEC colonization.



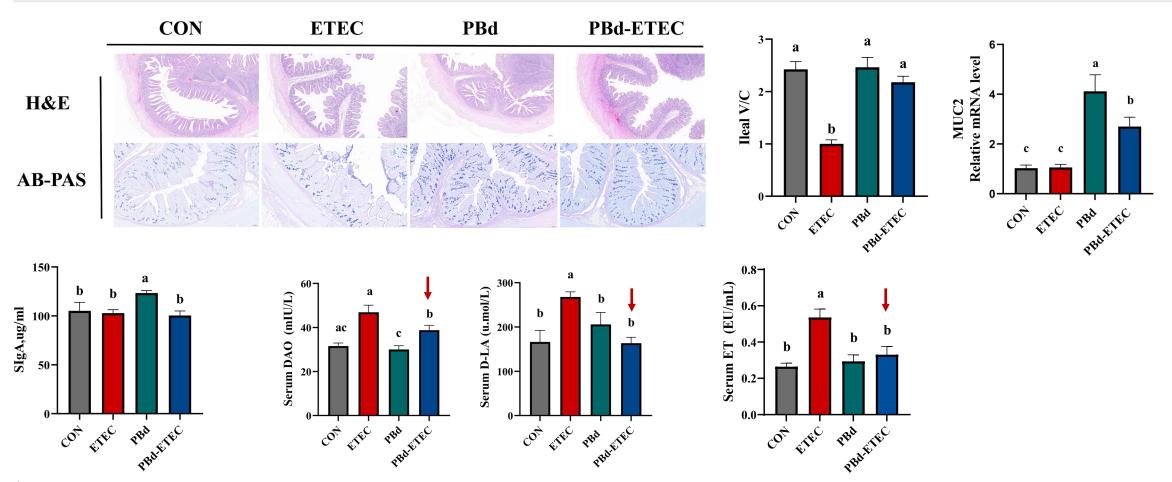
P. distasonis protected against ETEC-induced inflammation



✓ *P.distasonis* administration decreased the pro-inflammatory cytokines' expression and levels of ileum in ETEC-challenged piglets.

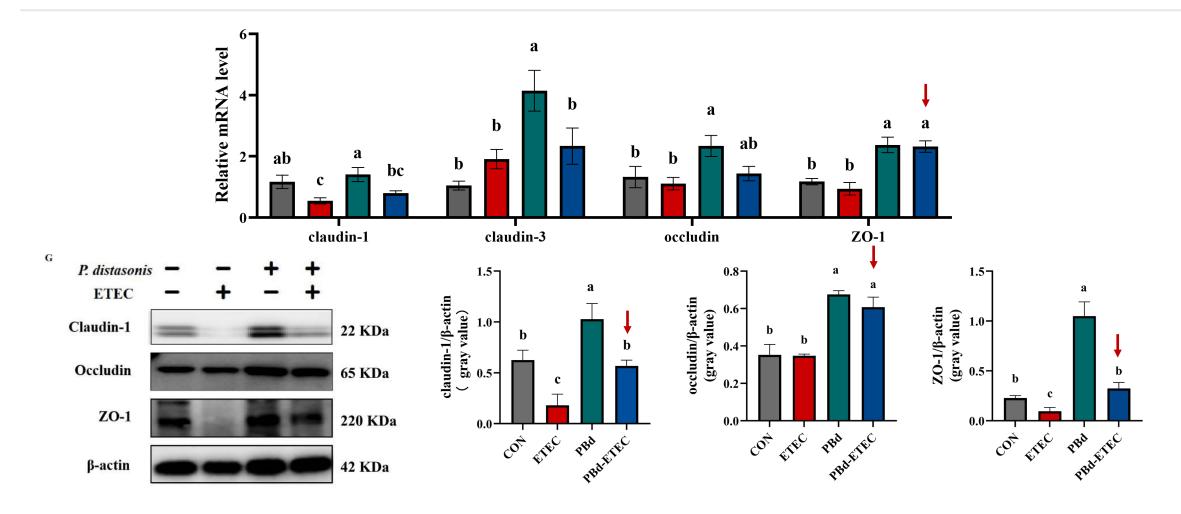


P. distasonis improved intestinal barrier integrity in piglet



- ✓ *P.distasonis* improved ileal barrier morphology, and increased Muc2 expression and sIgA level of piglets with or without ETEC infection.
- ✓ *P.distasonis* significantly decreased DAO, D-LA and ET levels in ETEC-challenged piglets.

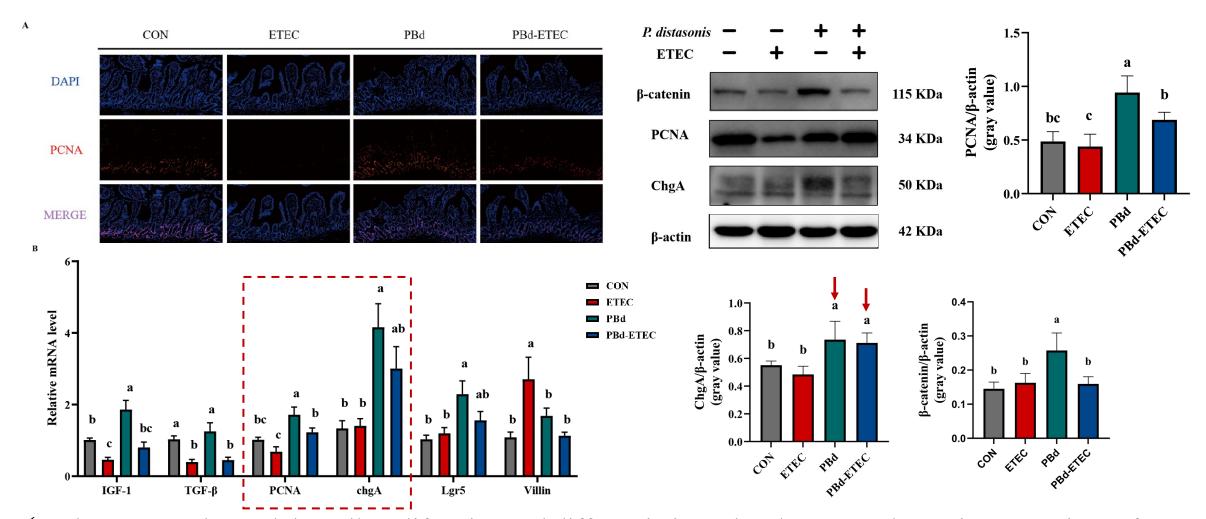
P. distasonis improved intestinal barrier integrity in piglet



✓ *P.distasonis* increased the intestinal tight junction mRNA expressions and protein abundances, which indicated that *P.distasonis* might block the invasion of enterotoxin by enhancing the integrity of intestinal barrier.



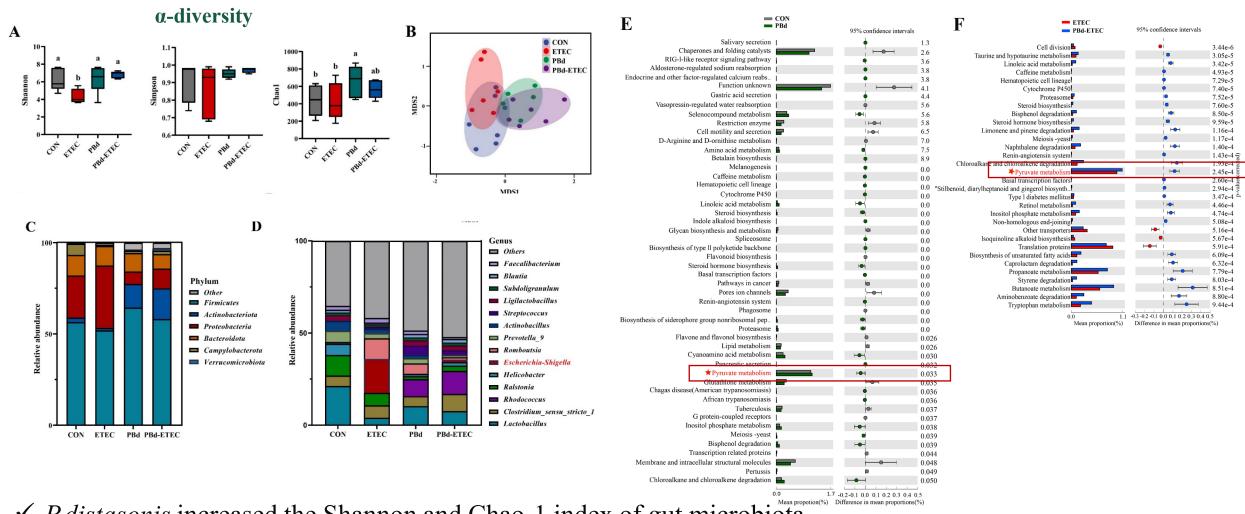
P. distasonis promoted intestinal development in piglet



✓ *P.distasonis* enhanced the cell proliferation and differentiation related genes and protein expressions of enterocyte in piglets with or without ETEC infection.



P. distasonis affected the gut microbial composition in piglets

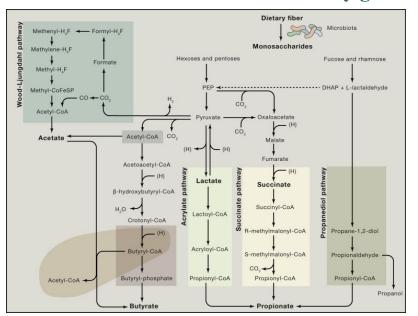


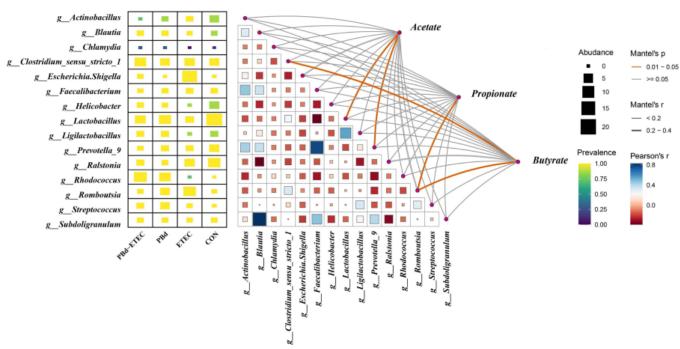
- ✓ *P.distasonis* increased the Shannon and Chao-1 index of gut microbiota.
- ✓ NMDS plot showed Pbd-ETEC group formed a distinct cluster separated from the ETEC group.
- ✓ KEGG analysis indicated *P.distasonis* might mainly participated in microbial pyruvate metabolism.



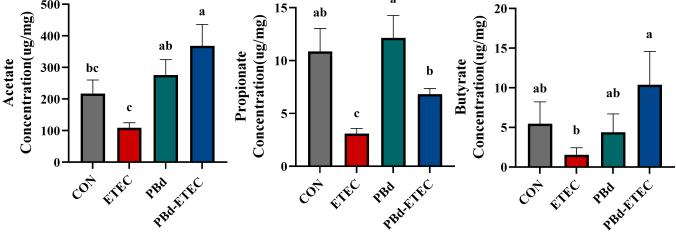
P. dastasonis raised SCFAs concentration of intestinal content

Pyruvate can be metabolized into SCFAs by gut microbes



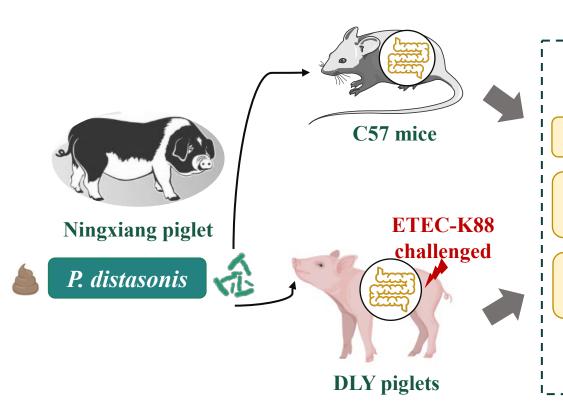


Mantel test



✓ *P.distasonis* might be involved in microbial SCFAs production.





Increase body weight gain

Improve intestinal barrier morphology and integrity

Promote intestinal development and growth

Decrease diarrhea rate

Inhibit ETEC colonization

Alleviate gut injury and inflammatory response

Optimize gut microbial composition

Promote SCFA production



Acknowledgement











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Thank You!