

FMD

Boehringer Ingelheim
The Veterinary Public Health Center

Claude HAMERS, 26 Oct 2017



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A MALAYSIA-12 PROTECTION AGAINST VIRULENT CHALLENGE: AN EXAMPLE OF CLINICAL PROTECTION OF A VACCINE DESPITE LOW R1 VALUES

The Veterinary Public Health Center
Dr. Claude HAMERS
26 Oct 2017



✓ FMDV is highly variable

- ✓ Need for monitoring of the circulating FMD viruses
- ✓ Need for vaccines having broad cross protection
- ✓ Need for tool allowing assessment of vaccine matching

✓ Vaccine matching r_1 values

- ✓ 0.3 threshold
- ✓ r_1 is quite variable

$$r_1 = \frac{\text{Ab titre of serum against field isolate (heterologous)}}{\text{Ab titre of serum against vaccine strain (homologous)}}$$

✓ Example based on 2 PD_{50} studies

✓ Low r_1 -values and high potency vaccines

- ✓ r_1 -value is part of a puzzle

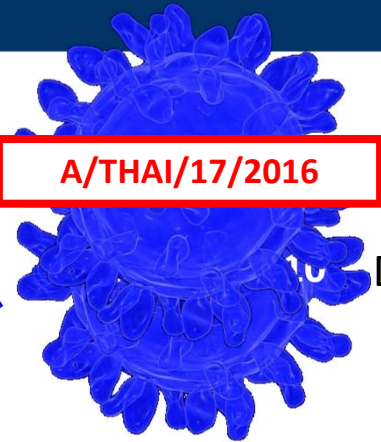
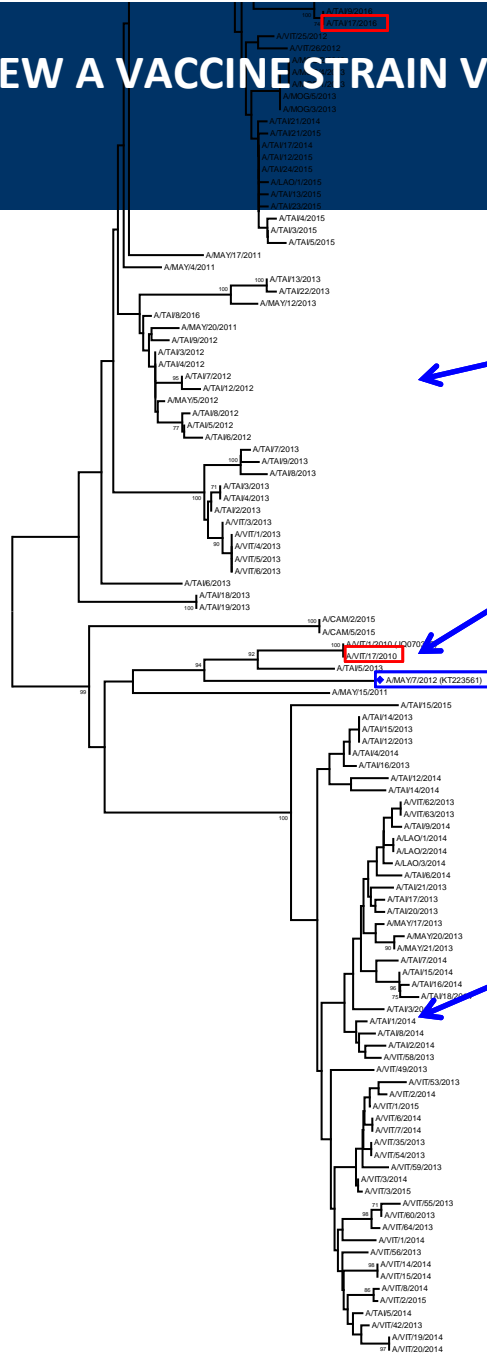
NEW A VACCINE STRAIN VS 2 DIFFERENT FIELD ISOLATES, *IN VIVO*



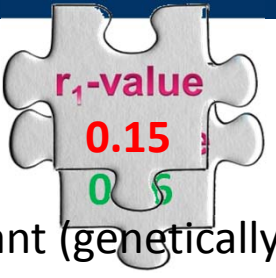
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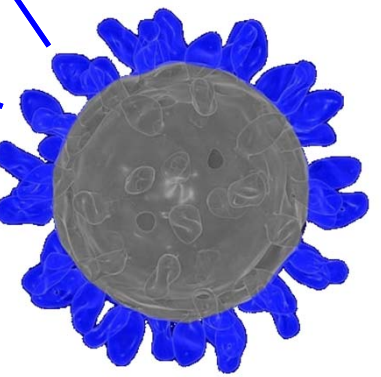
A/ASIA/Sea-97



A/THAI/17/2016



Distant (genetically and antigenically) and recent challenge strain
Closely related (genetically and antigenically) challenge strain



New Vaccine strain from A/ASIA/Sea-97 lineage, in monovalent DOE vaccine

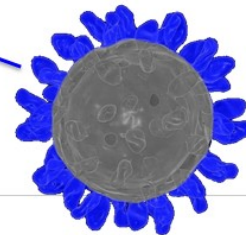
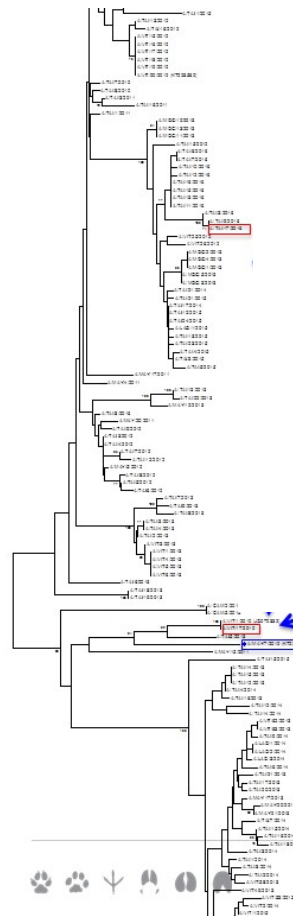
PD₅₀ STUDY N°1



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in vivo study



Closely related (**genetically** and **antigenically**) challenge strain

New Vaccine strain from
A/ASIA/Sea-97 lineage



Merial is now part of Boehringer Ingelheim

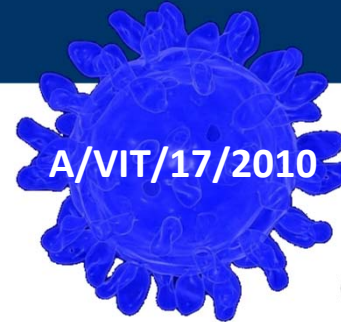


Study performed at WBVR, The Netherlands



New A/Sea-97 vaccine strain

Challenge at 21 dpv



A/VIT/17/2010



r_1 -value
0.56

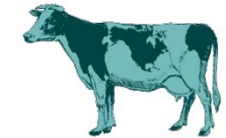
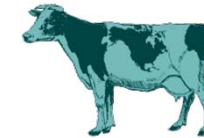
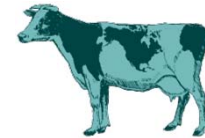
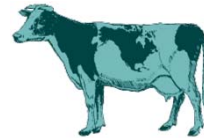
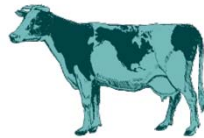
Closely related (genetically and antigenically) challenge strain

Monovalent DOE Vaccine

VACCINATION AT 3 DIFFERENT DOSES



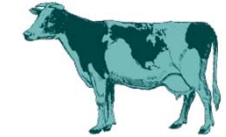
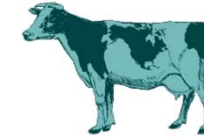
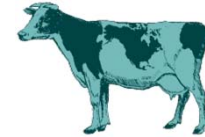
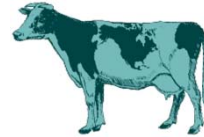
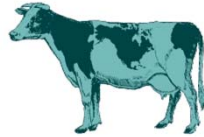
2 mL



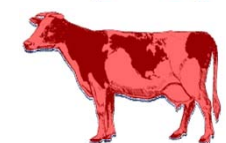
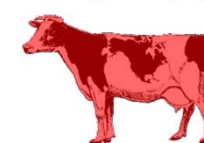
3 x 5 vaccinated cattle



0.5 mL



0.125 mL



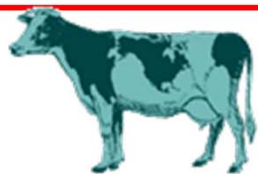
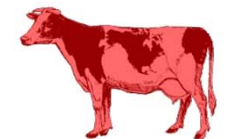
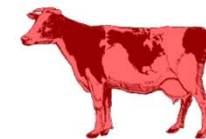
2 non-vaccinated cattle



Potency

> 18 PD₅₀/dose

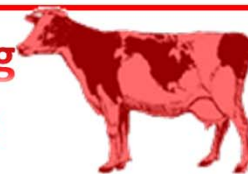
Controls



Protected Eff

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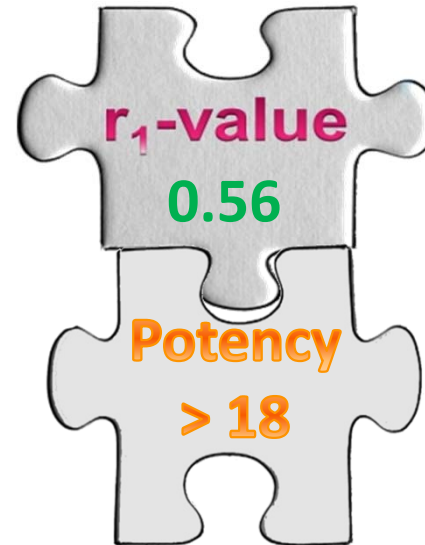
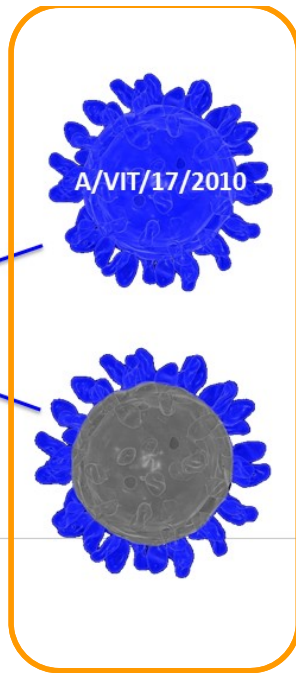
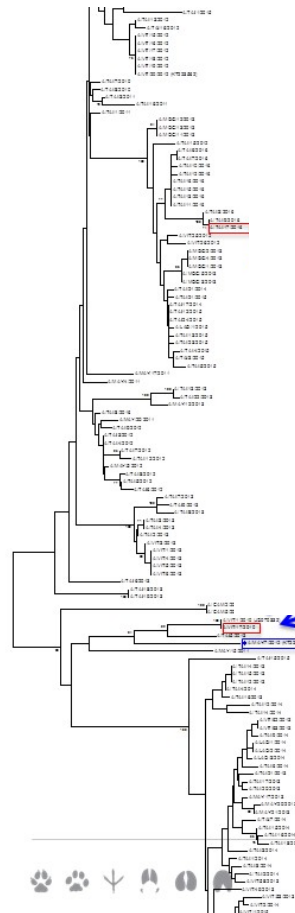


Not protected

**CONCLUSION:
NEW « A » VACCINE STRAIN IS HIGHLY POTENT (>18 PD₅₀)
AGAINST A CLOSELY RELATED ISOLATE**

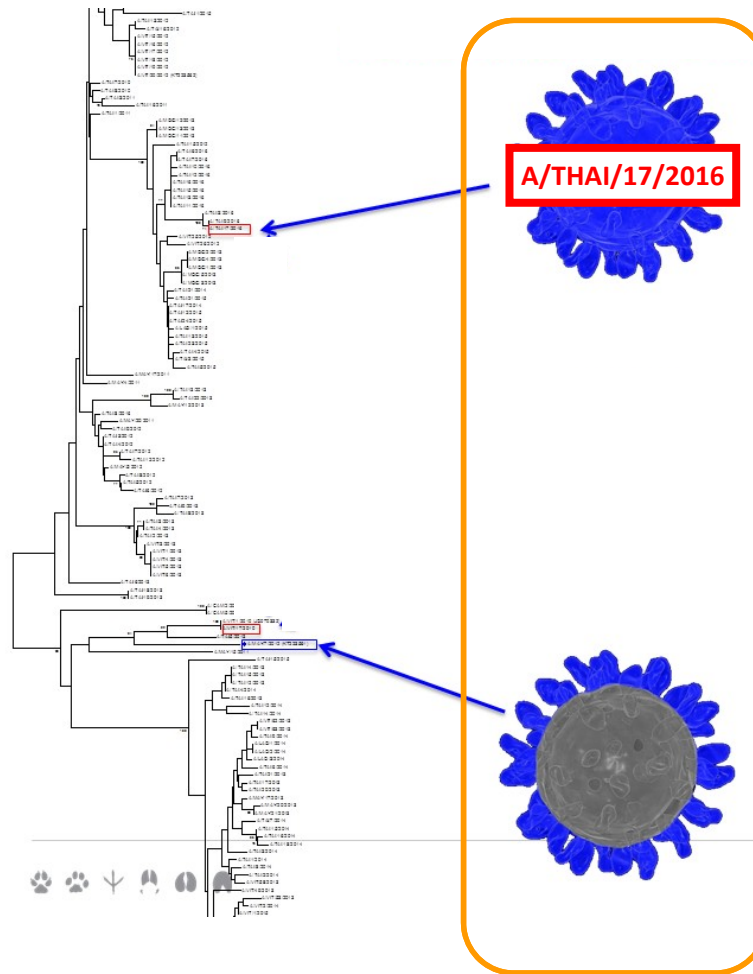


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PD₅₀ STUDY N°2

in vivo study



r_1 -value
0.15

Distant (**genetically and antigenically**) and recent challenge strain

New Vaccine strain from A/ASIA/Sea-97 lineage



Merial is now part of Boehringer Ingelheim

Study performed at TPI, UK



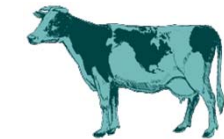
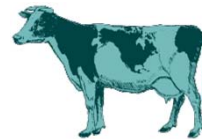
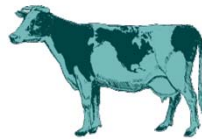
Challenge at 21 dpv



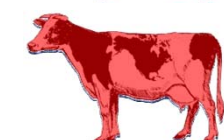
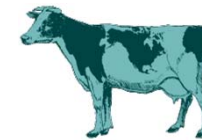
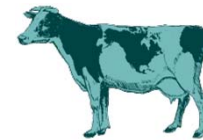
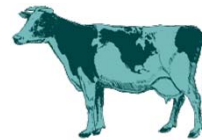
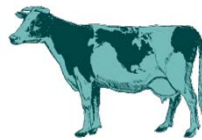
Monovalent DOE
Vaccine

Distant (genetically and antigenically)
and recent challenge strain

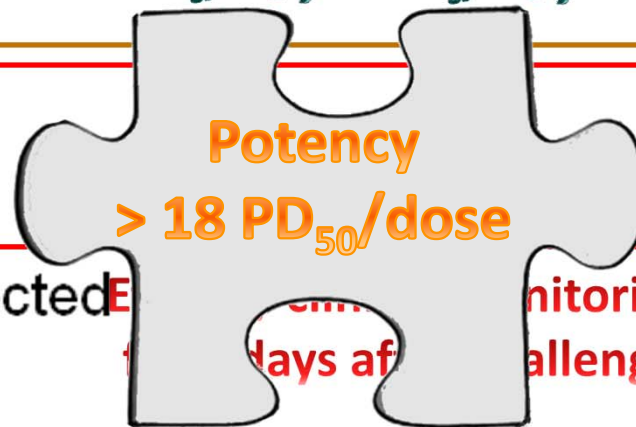
VACCINATION
AT 3 DIFFERENT DOSES



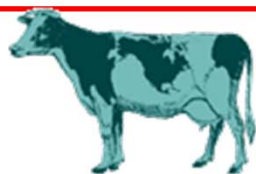
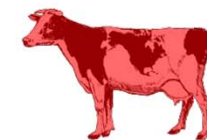
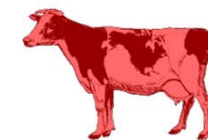
3 x 5 vaccinated cattle



2 non-vaccinated cattle



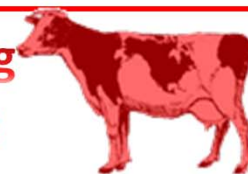
Controls



Protected

21 days af

Monitoring
challenge



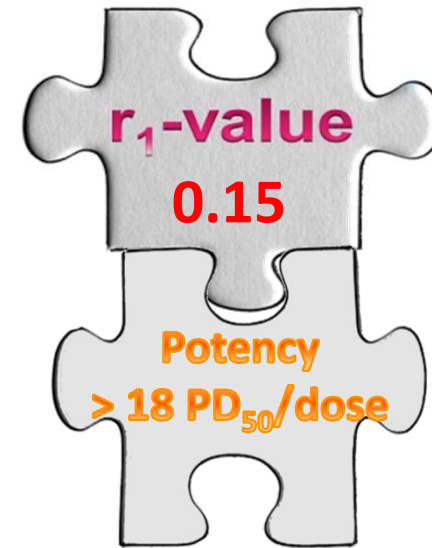
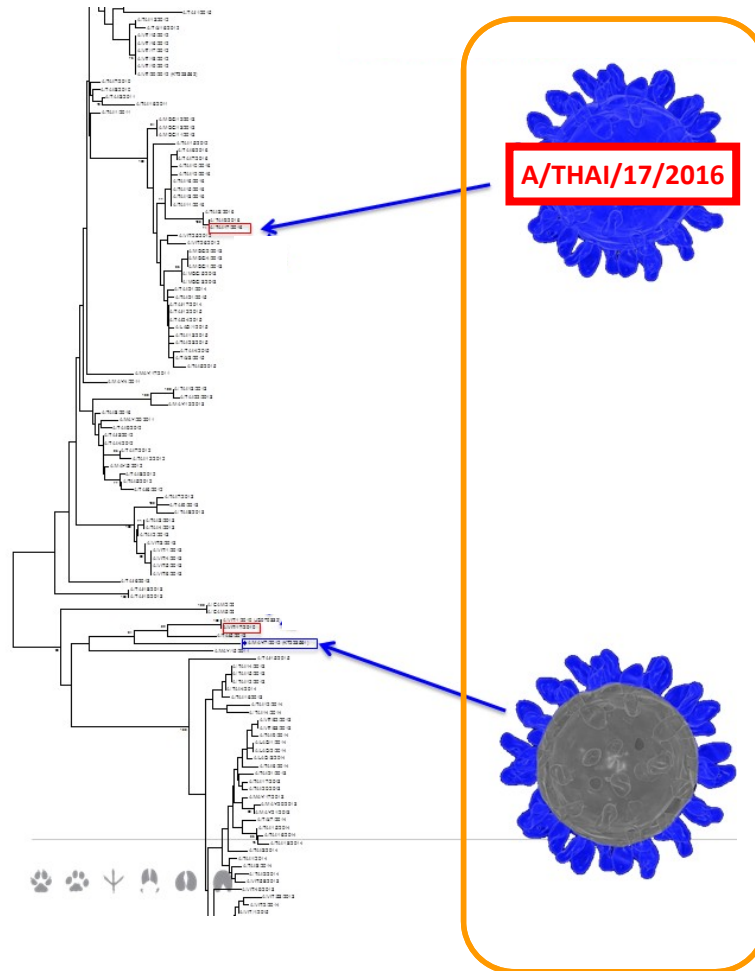
Not protected

CONCLUSION:

NEW « A » VACCINE STRAIN IS HIGHLY POTENT (>18 PD₅₀) AGAINST A RECENT & GENETICALLY/ANTIGENICALLY DISTANT ISOLATE



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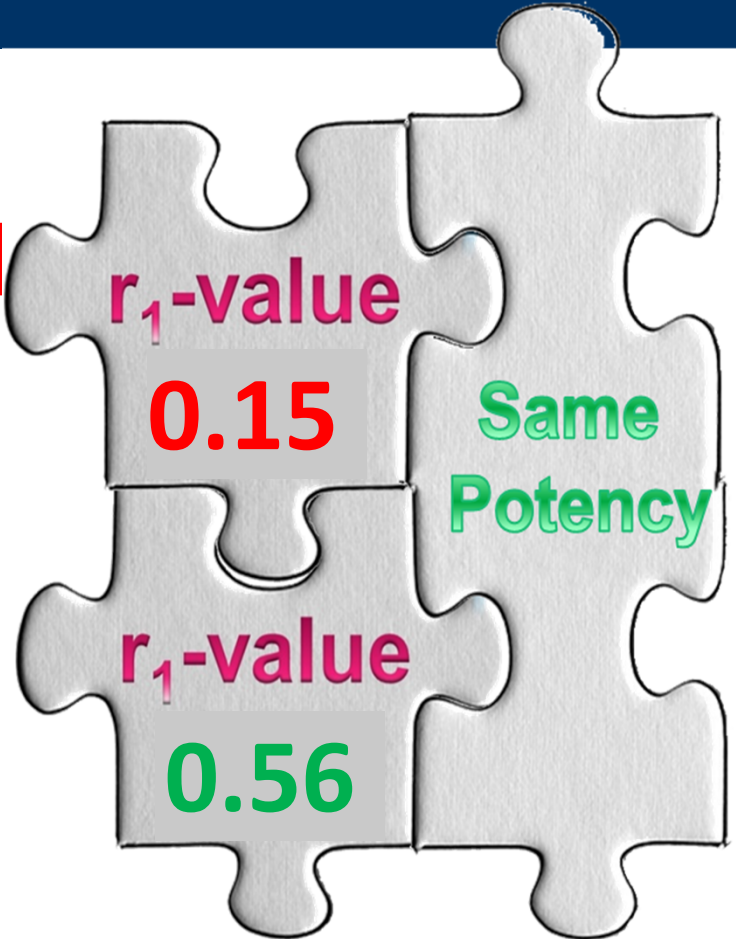
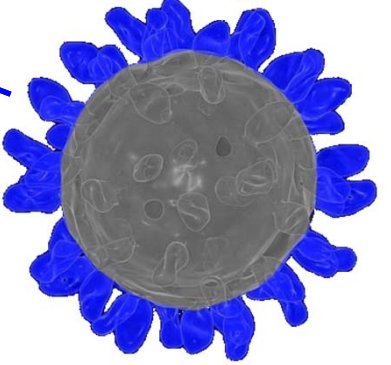
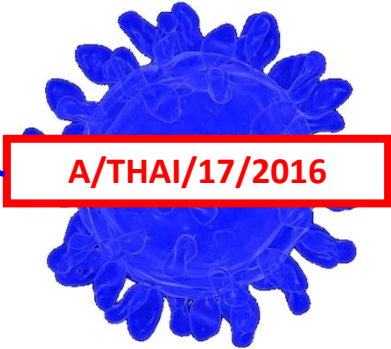
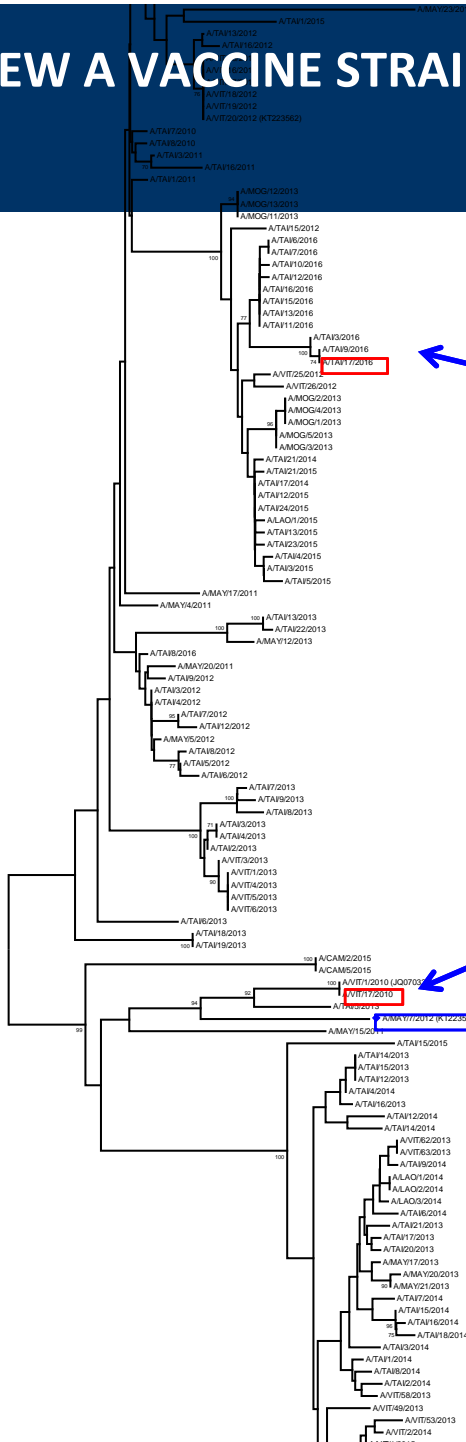
NEW A VACCINE STRAIN VS 2 DIFFERENT FIELD ISOLATES, *IN VIVO*



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A/ASIA/Sea-97



TAKE HOME MESSAGES

- ✓ **Now widely acknowledged that high potency vaccines are able to protect against antigenically distant strains, even in case of low r_1 -values**
- ✓ **Need for more robust method to assess heterologous vaccine performance**
- ✓ **Need to educate stakeholders on strengths/limitations of r_1 -values**
- ✓ **Success of a vaccination campaign does not solely depend on the antigenic match:**
 - Potency of the vaccine
 - Antigen quality and payload
 - Formulation (adjuvants)
 - Deployment of the vaccine, cold chain etc
 - Vaccination coverage and vaccination regimes
 - Interval between vaccination and exposure (DoI)
 - Severity of the challenge
 -



ACKNOWLEDGEMENTS

Wageningen Bioveterinary Research

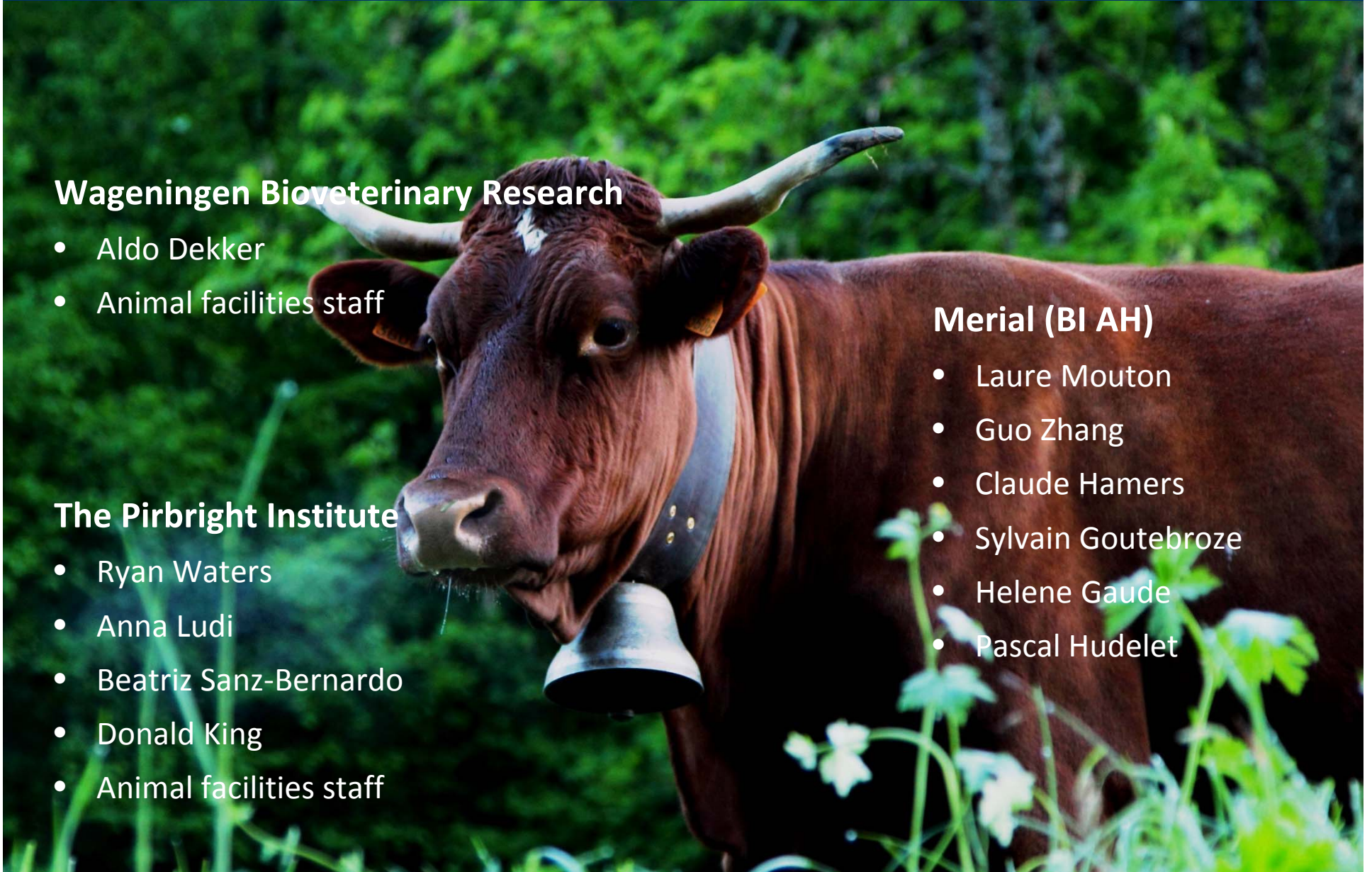
- Aldo Dekker
- Animal facilities staff

The Pirbright Institute

- Ryan Waters
- Anna Ludi
- Beatriz Sanz-Bernardo
- Donald King
- Animal facilities staff

Merial (BI AH)

- Laure Mouton
- Guo Zhang
- Claude Hamers
- Sylvain Goutebroze
- Helene Gaude
- Pascal Hudelet



THANK YOU FOR YOUR ATTENTION



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Introduction



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