



# Bienestar en Animales de Producción

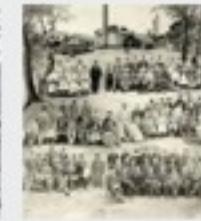
## Nuevas Oportunidades de Desarrollo

José M<sup>a</sup> Salleras Montells – Foro INIA 09.04.2013

# Boehringer Ingelheim compañía químico-farmacéutica fundada en 1885



## Principales fechas de nuestra historia



**1885**

Ernst Boehringer (1860-1892) adquiere en nombre de su hermano Albert una pequeña fábrica de ácido tartárico en Ingelheim, Alemania.

**1886**

Albert Boehringer inicia la producción de ácido tartárico.

**1893**

Durante los experimentos para producir ácido citrato, una fermentación inesperada ocasiona la formación de ácido láctico. En lugar de cancelar los experimentos, Albert Boehringer desarrolla el proceso con el propósito de producir ácido láctico a escala industrial.

**1895**

Da comienzo la fabricación industrial de ácido láctico, que en los siguientes años demostrará ser un éxito comercial.

**1907**

Se crea un fondo de beneficios para los empleados jubilados. Comienza la edificación de las primeras viviendas para empleados.

**1912**

Laudanón® se convierte en la primera especialidad farmacéutica de la compañía.

**1935**

50º aniversario. La compañía emplea entonces a 865 personas.



**1951**

Lanzamiento de Buscopan®, analgésico y espasmolítico.

**1955**

La compañía crea la división de Salud Animal.

**1973**

Se funda Boehringer Ingelheim GmbH como holding alemán de las filiales extranjeras del grupo.

**1985**

Boehringer Ingelheim celebra su centenario con 22.250 empleados, 8.784 de ellos en Alemania.

**1999**

Lanzamiento de Micardis®, medicamento antihipertensivo.

**2002**

Lanzamiento de Spiriva®, fármaco empleado en el tratamiento de la EPOC.

**2008**

Lanzamiento de Pradaxa®, un anticoagulante oral.

**2010**

Boehringer Ingelheim celebra su 125º aniversario.

## Breve recorrido por la historia

**1952**

Compra de Laboratorios Barry.  
Cambio de nombre a C.H. Boehringer Sohn  
Ingelheim S.A.E.  
Traslado a Barcelona.  
Compra de Laboratorios Rhei, S.A. en  
Barcelona.

**1960**

Se inaugura la planta química en Mataró  
de Mar con la extracción de alcaloides a  
partir de productos naturales.

**1963**

Inauguración de la planta de producción  
farmacéutica en Sant Joan Despí.

**1970**

Adquisición de Europharma, S.A. (Madrid).

**1972**

Se crea el Área de Veterinaria.

**1998**

Traslado de la planta de producción  
farmacéutica a Sant Cugat del Vallès.

**1999**

Inauguración oficial de la planta de Sant  
Cugat del Vallès a cargo de S.S. MM. los  
Reyes de España.

**2001**

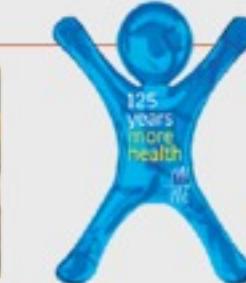
Traslado de la sede social y administrativa  
de Barcelona a Sant Cugat del Vallès.

**2002**

Celebración del 50 aniversario de  
Boehringer Ingelheim España.

**2008**

30º aniversario de la planta de producción  
farmacéutica de Sant Cugat del Vallès.

**2010**

Celebración del 50 aniversario de la planta  
química de Mataró de Mar.

Celebración del 125 aniversario del  
nacimiento de Boehringer Ingelheim.

**2012**

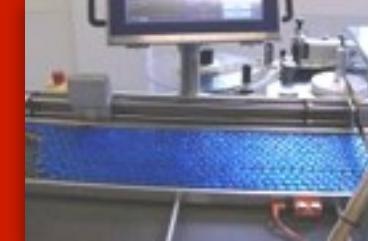
Celebración del 40 aniversario del Área de  
Veterinaria.

## Biofarmacéutica

I+D+i



Fabricación

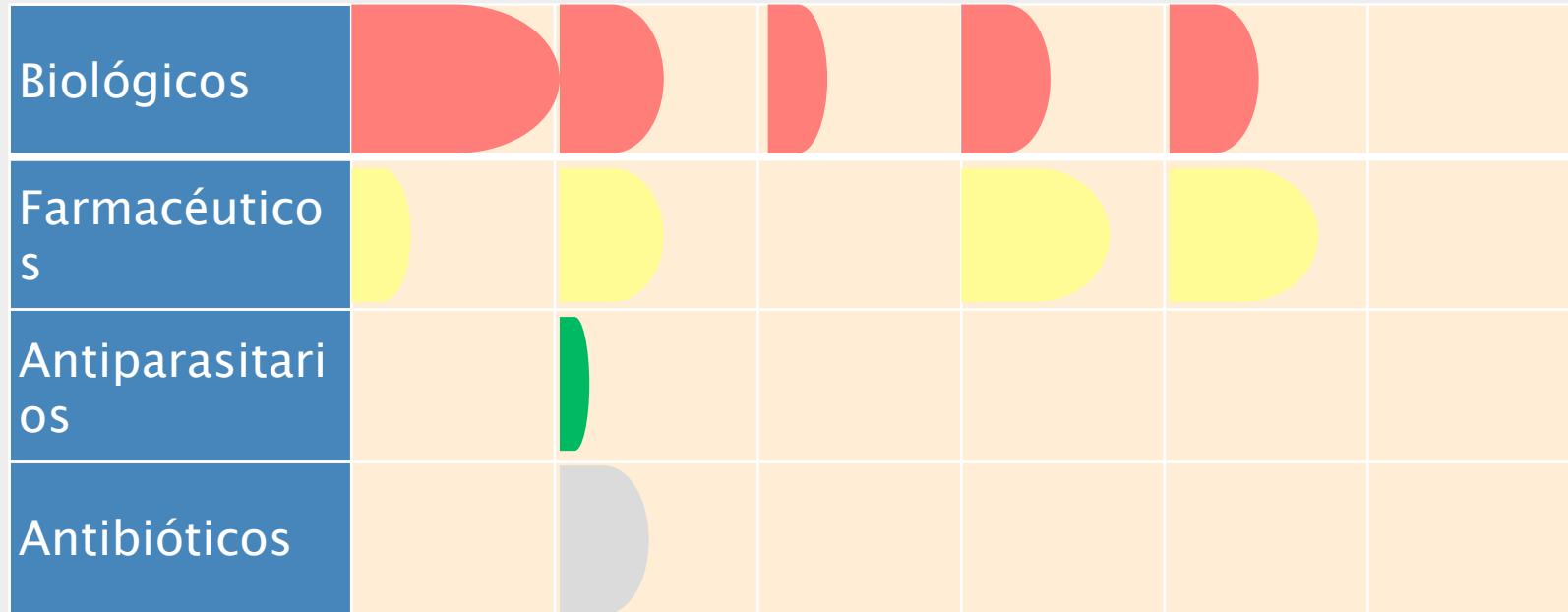


Marketing

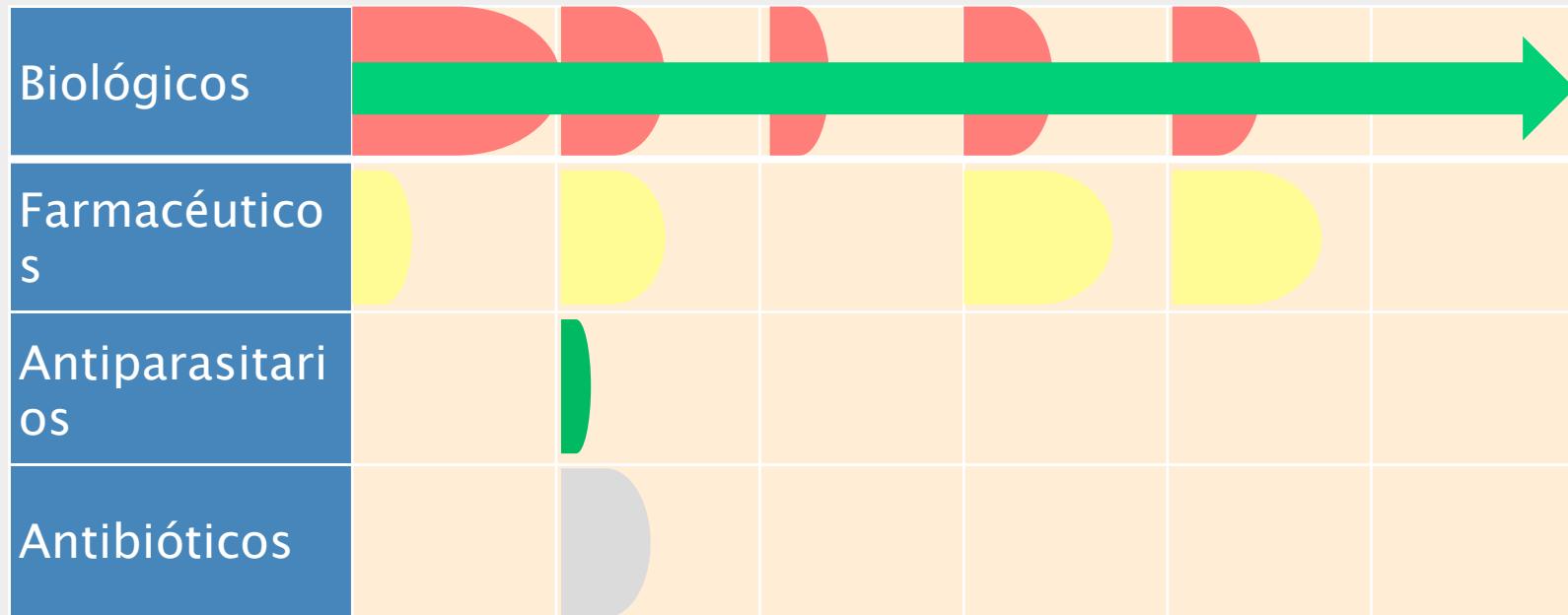


Aportar valor a través de la innovación

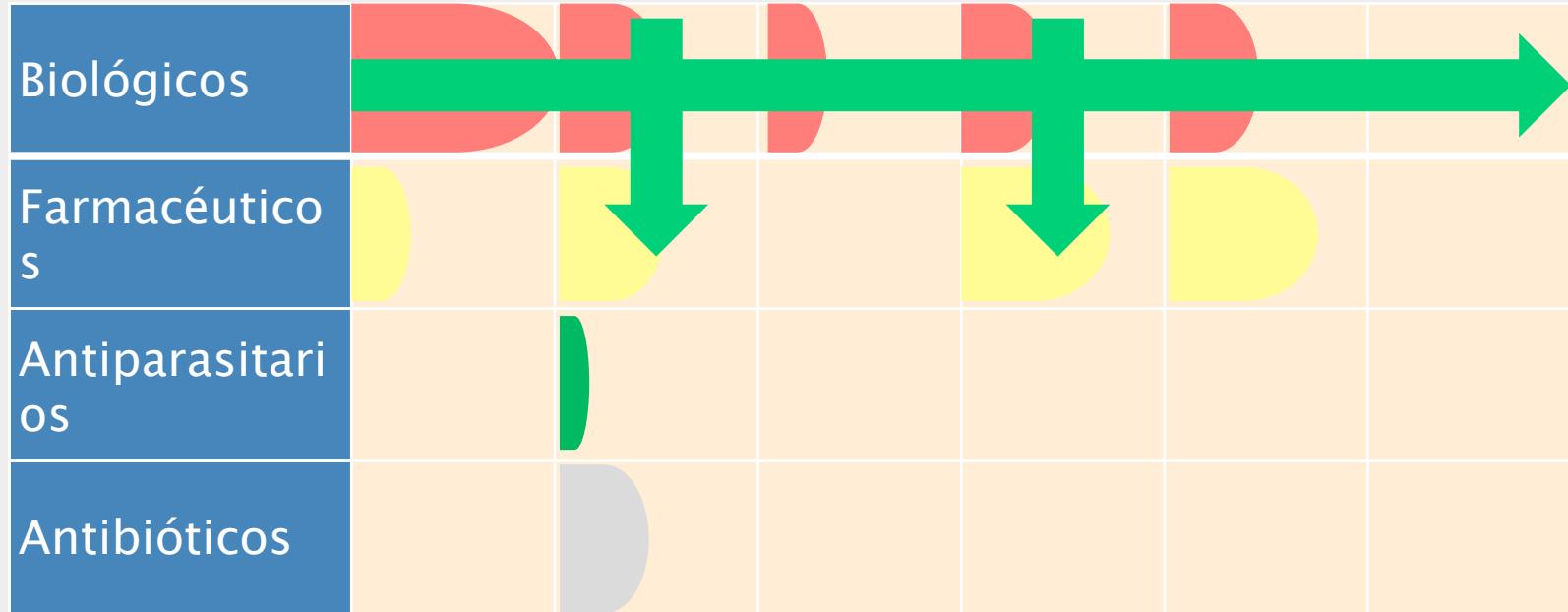
# Portfolio Actual y Segmentos de Desarrollo animal health



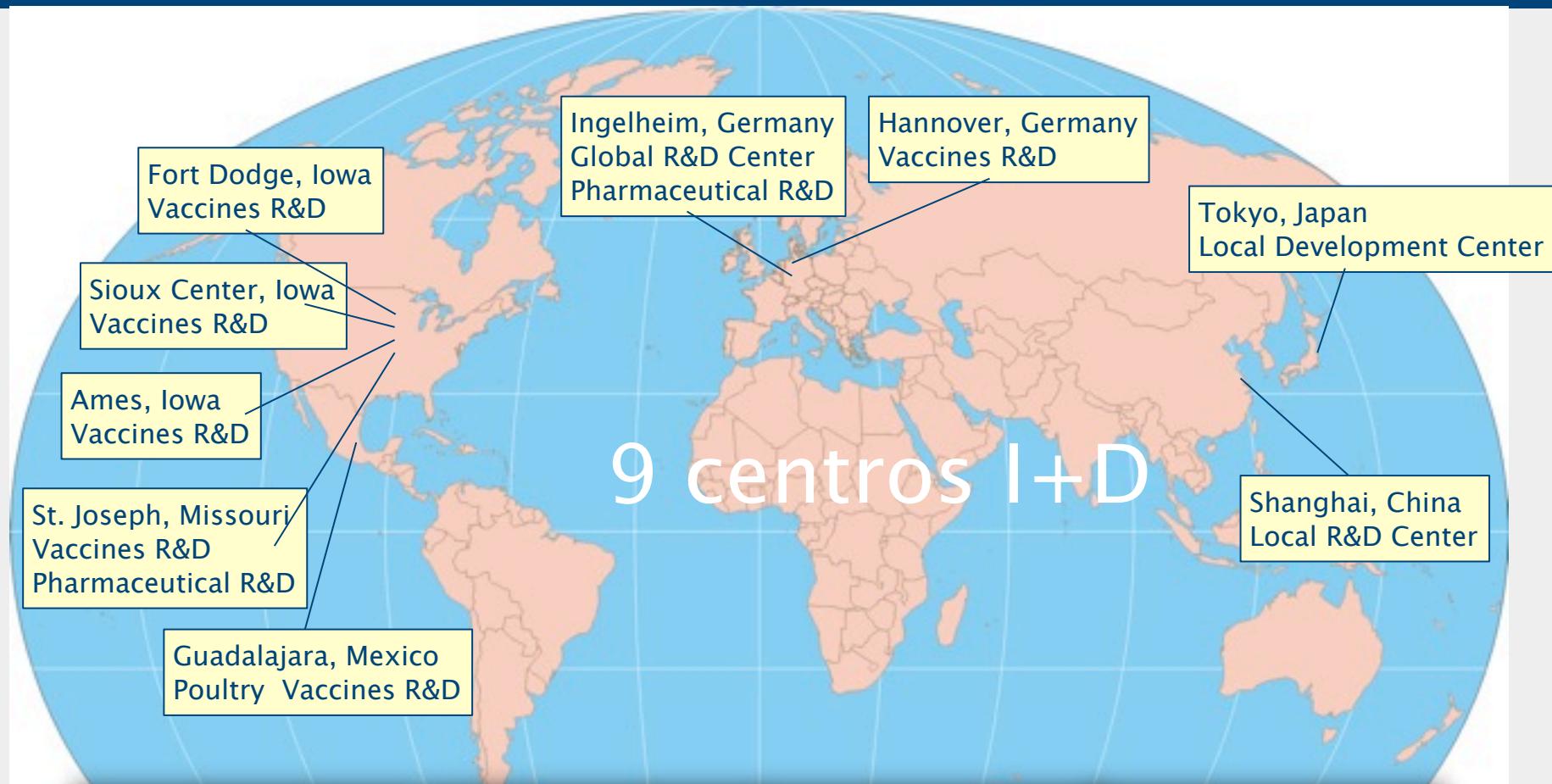
# Portfolio Actual y Segmentos de Desarrollo animal health



# Portfolio Actual y Segmentos de Desarrollo animal health



# Centros de Investigación y Desarrollo animal health



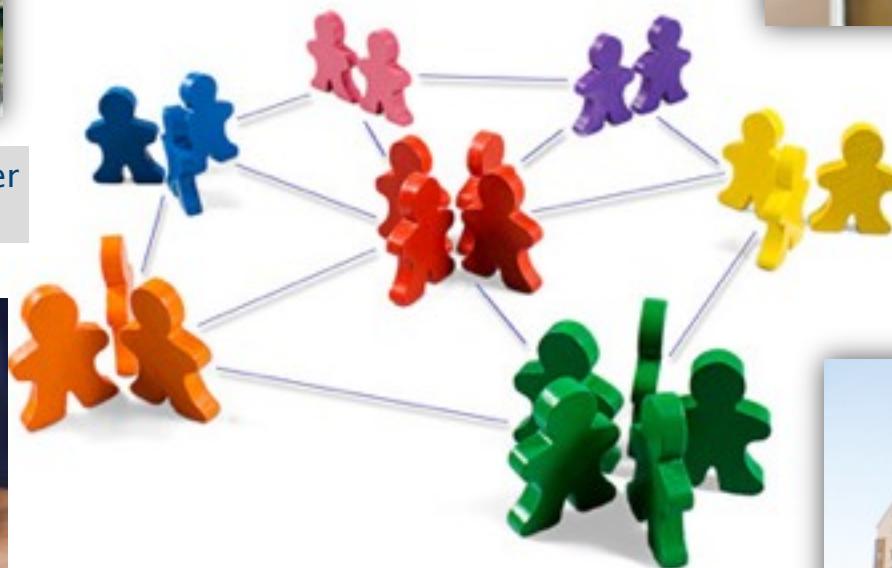
# Alianzas y Colaboraciones con equipos de investigación a nivel mundial



# Bienestar Animal compromiso y responsabilidad

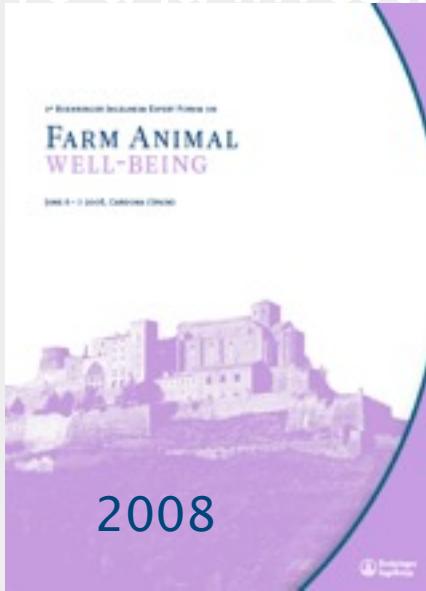


Vaccines R&D Research Center  
Hannover, Germany

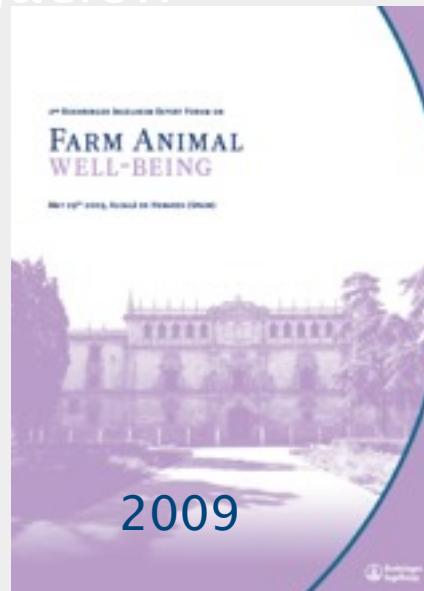


Vaccines R&D Research Center  
St. Joseph, Missouri

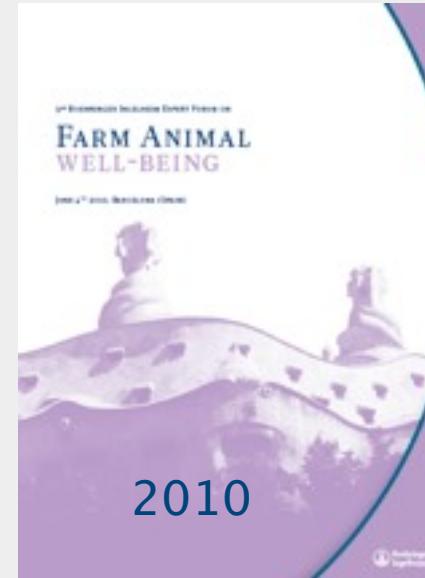
# Foro Internacional Bienestar Animales Producción apoyo a la investigación



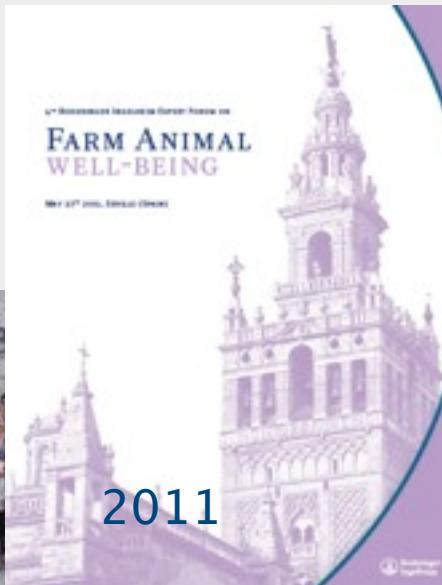
2008



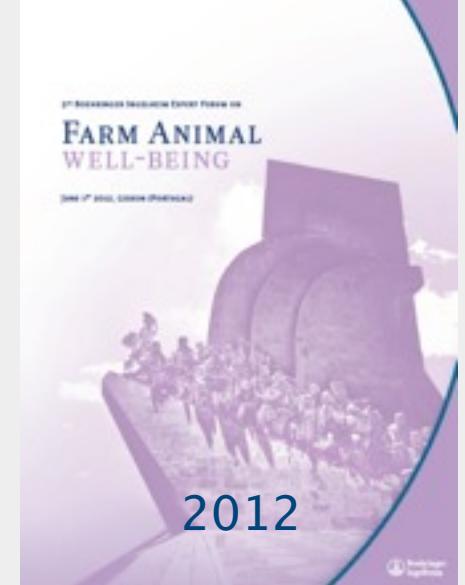
2009



2010



2011



2012

# Foro Internacional Bienestar Animales

## Producción

### apoyo a la investigación



2011

2012

# Farm Animal Welfare Education Center apoyo a la formación



## Farm Animal Welfare Education Centre

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- [Cursos de Formación](#)
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FAWEC (Farm Animal Welfare Education Centre – Centro de Educación en Bienestar de Animales de Producción) ha sido creado por el Servicio de Nutrición y Bienestar Animal (SNIBA) del Departamento de Ciencia Animal y de los Alimentos de la Facultad de Veterinaria de la Universitat Autònoma de Barcelona (UAB).

El Centro tiene como objetivo principal la formación en bienestar de animales de producción, mediante la publicación periódica de fichas técnicas que resumen los conocimientos actuales en bienestar animal y la organización de cursos teórico-prácticos de formación sobre el bienestar del vacuno de leche y del porcino.



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con el patrocinio de



# Oportunidades de Desarrollo productos más seguros ...



## Comparative safety of PCV2 vaccines under field conditions

John Kilk, DVM; Edgar Diaz, DVM  
Boehringer Ingelheim Veterinary, Inc., St Joseph, Missouri

### Introduction

Porcine Circovirus type 2 (PCV2) vaccines have been reported to be highly effective at reducing mortality due to Porcine Circovirus Associated Disease (PCVAD).<sup>1</sup> However, several of them may also induce significant lesions following injection. In one Canadian study, over 16% of pigs (5,900 of 35,108 pigs) developed injection site lesions up to 3cm in diameter following injection according to label directions.<sup>2</sup> Reactivity of certain PCV2 vaccines has also been reported from the United States.<sup>3</sup> In a laboratory experiment conducted as a pilot project to assess the comparative severity of lesions induced by various commercially available PCV2 vaccines, the Fort Dodge and Intervet PCV2 vaccines were found to induce significantly more severe lesions than saline control injection sites and Boehringer Ingelheim (BI) PCV2 vaccine. BI vaccine injection site lesion scores were not significantly different from saline controls.<sup>4,5</sup> This second study was conducted to compare the same vaccines under field conditions.

### Materials and methods

This field study compared the safety of two commercial vaccines under field conditions. Two groups of three treatment groups were used. Each group consisted of the various vaccination sites on the side of the neck at the level of the last cervical vertebrae. One dose of Ingelvac® MycoFLEX™ or Ingelvac® CircoFLEX™ PCV2 One Dose vaccine (2mL) was administered. Pigs received a second dose of vaccine 14 days after the first injection, i.e., a total of two doses of equal volume were injected into the side of the neck to serve as a comparison. Each dose was made with multiple needles per needle. Pigs were randomly assigned to each treatment group and were injected while standing.

A subset of twenty pigs from each group were examined at 14 days post-vaccination. These pigs were palpated and examined grossly prior to humane

euthanasia and injection sites were harvested and preserved in formalin. Histological lesions were blindly scored at the Iowa State University Veterinary Diagnostic Laboratory on a 1-5 scale.<sup>3</sup>

The null hypothesis was there would be no differences in post-vaccination reactivity or injection site lesion scores between treatments. Statistical analyses were made on normal, continuously distributed variables by ANOVA with Tukey's HSD utilized to identify which groups were significantly different from one another (JMP; Cary, North Carolina). Student's t-tests were used for pair-wise comparisons (JMP; Cary, North Carolina). Proportions were analyzed by Pearson's Chi Square tests (Statistica 8.0; Tallahassee, Florida).

### Results and discussion

Five of 300 (5/300) pigs died within the first 24 hours post-injection in the Fort Dodge vaccine group, with the deaths attributed to anaphylactic reaction to vaccine. No pigs died during the study period in the Ingelvac vaccine groups.

Injection site lesions were examined in all pigs.

Injection site lesions were compared to saline injection site lesions. There was no difference in lesion scores of Ingelvac MycoFLEX versus Ingelvac CircoFLEX (0.048 ± 0.218 versus 0.048 ± 0.218, respectively). The most severe lesion was observed in a pig that received two doses of Ingelvac MycoFLEX versus Ingelvac CircoFLEX (1.095 ± 0.301 versus 1.095 ± 0.301, respectively).

Injection site lesions were compared to the resulting injection site lesion scores. Ingelvac CircoFLEX induced a greater number of injection site lesions than Ingelvac MycoFLEX (1.71 vs 0.57,  $P < 0.0001$ , Figure 2). The



## An innovative method for quantifying animal behavior responses to various immunization protocols

Keith Bresee; Roy Edler; Edgar Diaz  
Boehringer Ingelheim Veterinary, Inc., St Joseph, Missouri

### Introduction

Pigs are naturally curious about their environment and seek out any recent changes or introductions into that environment. Pigs are also curious, initially acting fearful or excitable when a person first enters a pen. However, after a 15 second adjustment period, pigs that have no previous reason to fear people will relax and explore the presence of a person in the pen by nuzzling or biting at the observer's legs and feet. The National Pork Board's Swine Welfare Assurance Program<sup>®</sup> (SWAP) introduced in 2003 utilized these behavioral principles to identify and quantify pigs that are relaxed in the presence of people in a pen. After entering a pen and knowing down in that pen, 50% of the pigs in that pen should return to the observer's gloved hand or approach the observer and/or demonstrate a relaxed posture within the 15 seconds of the animal observing the observer. This has been employed as a welfare assessment approach may be used to evaluate the relative reactivity (sensitivity) of the animal to a stimulus.

A similar adaptation of the 2003 SWAP protocol was used to evaluate the behavior of pigs in pens containing pigs from the Iberian/Schroeder<sup>®</sup> 1/2 dose combination Ingelvac® MycoFLEX<sup>®</sup>, a 1/2 dose Ingelvac® MycoFLEX<sup>®</sup>, a 1/2 dose Ingelvac® CircoFLEX<sup>®</sup>, a 1/2 dose Ingelvac® CircoFLEX<sup>®</sup>, a 1/2 dose Ingelvac® MycoFLEX<sup>®</sup> and a 1/2 dose Ingelvac® CircoFLEX<sup>®</sup>. All pigs were infected with circovirus disease virus. Ingelvac® MycoFLEX and Ingelvac® CircoFLEX are both licensed for use in pigs and are caused by Mycoplasma suis and circovirus disease, respectively. Ingelvac® MycoFLEX<sup>®</sup> because it was the first product to be licensed in the United States. This study evaluated the relative reactivity of pigs to the Ingelvac® MycoFLEX<sup>®</sup> and Ingelvac® CircoFLEX<sup>®</sup> vaccines. This evaluation was conducted by mixing/combining Ingelvac® MycoFLEX<sup>®</sup> and Ingelvac® CircoFLEX<sup>®</sup> in a 1:1 ratio. A subset of twenty pigs were used in this study. The pigs were randomly assigned to receive either Ingelvac® MycoFLEX<sup>®</sup> or Ingelvac® CircoFLEX<sup>®</sup>. The pigs were palpated and examined grossly prior to humane euthanasia and injection sites were harvested and preserved in formalin. Histological lesions were blindly scored at the Iowa State University Veterinary Diagnostic Laboratory on a 1-5 scale.<sup>3</sup>

One dose was not quantified. The terms "safety", "reactivity", and "approachability" as used in this document refer to relative responses by groups of pigs to various injectable vaccines or saline as measured by the relative decrease in percent of pigs in a group that are willing to approach an observer within a specified time interval, with the baseline value for each group determined prior to treatment and the same measurement repeated 6 hours post-treatment.

### Objectives

To evaluate the comparative reactivity of Ingelvac® MycoFLEX alone, Ingelvac® MycoFLEX alone, Ingelvac® CircoFLEX/Ingelvac® MycoFLEX mixture, Circovac®/M+PAC mixture, and saline injection as measured by clinical observations pre-injection and 6 hours post



# Oportunidades de Desarrollo combinaciones de productos ...



## Efficacy evaluation of a mixed *Mycoplasma hyopneumoniae* bacterin and a porcine circovirus type 2 vaccine

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Philip W. Hayes<sup>1</sup>, DVM; Ryan Salmonau<sup>2</sup>, DVM

<sup>1</sup>Boehringer Ingelheim Veterinary, Inc., St. Joseph, Missouri, <sup>2</sup>Veterinary Resources, Inc., Cambridge, Iowa

### Introduction

Diseases associated with *Mycoplasma hyopneumoniae* (*M. hyo*, enzootic pneumonia) and Porcine Circovirus Type 2 (PCV2) infections are a major concern in the swine industry. *M. hyo* has also been implicated as one of the major co-factors in the development of PCVAD.<sup>1,2</sup> The objective of these studies was to evaluate the efficacy of both *Mycoplasma hyopneumoniae* and Porcine Circovirus Type 2 vaccines when the monovalent USDA licensed vaccines for the two agents are mixed and administered in a single combined injection.

### Materials and methods

The efficacy of each fraction was evaluated in separate studies. This allowed for evaluation of *M. hyo* and PCV2 vaccine protection in their appropriate host animal challenge models.

#### *M. hyo* evaluation

The *M. hyo* efficacy evaluation was performed in conventional pigs approximately 3 weeks of age. Pigs were

### PCV2 evaluation

The PCV2 efficacy evaluation was performed in caesarian-derived, colostrum-deprived piglets. At approximately three weeks of age, piglets were vaccinated with Ingelvac MycoFLEX and Ingelvac CircoFLEX in a single 2 mL dose (mixed as described above). On day 31 post-vaccination, vaccinated and control animals were administered a virulent PCV2 challenge virus. Twenty two days following the administration of the challenge material, all animals were euthanized and selected tissues were removed and submitted for evaluation by histopathology and PCV2 immunohistochemistry (IHC). The primary efficacy criteria were prevalence of lymphoid depletion, lymphoid inflammation and lymphoid IHC. Tests for differences in percentages between treatment groups were performed using Fisher's Exact Test.

### Results

There were no systemic or injection site adverse reactions that could be attributed to the vaccine mixture in either study.

## Ingelvac MycoFLEX® + Ingelvac CircoFLEX®

Ingelvac MycoFLEX® Cada dosis de 1 ml de la vacuna inactivada contiene: *Mycoplasma hyopneumoniae* a 1 PR% \*Potencia Relativa (test/ELISA) / por comparación con una vacuna de referencia. Adyuvante: Carbómero. Indicaciones: Inmunización activa de cerdos, a partir de 3 semanas de edad, para reducir lesiones pulmonares después de una infección con *Mycoplasma hyopneumoniae*. Ínicio de la inmunidad: 2 semanas tras la vacunación. Duración de la inmunidad: al menos 26 semanas. Interacción con otros medicamentos: Esta información sobre la seguridad y la eficacia que demuestra que esta vacuna se puede mezclar con Ingelvac CircoFLEX de Boehringer Ingelheim y administrar en un punto de inyección. Reacciones adversas: Las reacciones adversas no tan comunes pueden observarse en el lugar de inyección una hinchazón transitoria de hasta 1-continente de diámetro, algunas veces asociada con eritema de la piel. Estas hinchazones pueden durar hasta 5 días. Puede observarse un incremento transitorio en la temperatura rectal de alrededor de 0,5 °C de promedio que dura hasta 24 horas después de la vacunación. Patología: Inyección única por vía intramuscular (IM) de una dosis (1 mL), preferiblemente en el cuarto de cerdos a partir de 3 semanas de edad. Tiempo de espera: Cerca de 3 semanas. Conservación: Conservar y transportar refrigerado (entre 2 °C y 8 °C). No congelar. Proteger de la luz. Usar inmediatamente después de abierta. Presentación: Frascos de 50 ml (50 dosis), 100 ml (100 dosis) y 250 ml (250 dosis). Registro n°: 2.034-ES2 Titular de la autorización: Boehringer Ingelheim Veterinaria GmbH, Ingelheim/Rhein, Alemania.

Ingelvac CircoFLEX® Cada dosis de 1 ml de la vacuna inactivada contiene: Proteína OPI2 de Circovirus Porcino Tipo 2, PR mínima 1,0; PR máxima 3,75. \*Potencia Relativa (test/ELISA) por comparación con una vacuna de referencia. Adyuvante: Carbómero. Indicaciones: Inmunización activa de cerdos de más de 2 semanas frente al PCV2 para la reducción de la mortalidad, signos clínicos- incluyendo pérdida de peso y lesiones en tejidos blandos, relacionadas con las enfermedades asociadas al PCV2 (PCVAD). Además, la vacunación ha demostrado reducir la excreción nasal de PCV2, la cospuricación en sangre y tejidos blandos y la duración de la elevación. Ínicio de la protección: 2 semanas tras la vacunación. Duración de la protección: al menos 17 semanas. Interacción con otros medicamentos: Esta información sobre la seguridad y la eficacia que demuestra que esta vacuna se puede mezclar con Ingelvac MycoFLEX® de Boehringer Ingelheim y administrar en un punto de inyección. Reacciones adversas: De forma muy frecuente se produce hinchazón, lejía y eritema el día de la vacunación. En muy raras ocasiones, pueden ocurrir reacciones anafilácticas que deberán tratarse sistemáticamente. Patología: Inyección única por vía intramuscular de una dosis (1 mL), independientemente del peso vivo. Tiempo de espera: Cerca de 3 semanas. Conservación: Conservar y transportar refrigerado (entre 2 °C y 8 °C). No congelar. Proteger de la luz. Usar inmediatamente después de abierto. Presentación: Frascos de 50ml (50 dosis), 100ml (100 dosis) y 250ml (250 dosis). Registro n°: 1.012.031.079/002/000-000/000-000/000 Titular de la autorización: Boehringer Ingelheim Veterinaria GmbH, Ingelheim/Rhein, Alemania.

# Oportunidades de Desarrollo nuevas indicaciones terapéuticas ...



Veterinary Anesthesia and Analgesia  
Formerly the Journal of Veterinary Anesthesia  
Veterinary Anesthesia and Analgesia, 2010, 30, 167-174  
doi:10.1111/j.1467-2993.2010.00546.x

## RESEARCH PAPER

### Pre-emptive meloxicam for postoperative analgesia in piglets undergoing surgical castration

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## Abstract

**Objective** To investigate the effect of preoperative meloxicam administration on postoperative stress and pain induced by surgical castration in piglets.

**Study design** Prospective, blinded, randomised clinical trial.

**Animals** One hundred and eighty male <1 week of age.

**Methods** Castration was performed on 100 piglets which had received either an intramuscular injection of 0.4 mg kg<sup>-1</sup> meloxicam or a placebo 30 minutes before the procedure. Blood cortisol and ACTH concentrations were determined at 0, 10 and 30 minutes post-castration and haptoglobin was measured at 24 hours post-castration. Presence of foreleg movements, hind-leg movements, faeces emission, tremors or other body movements were recorded during the castration procedure. Scores for presence or absence of prostrations, tail movements and isolation were recorded at 0, 10, 30 minutes, and at 1, 2, 4 and 24 hours post-castration and combined in a global behavioral score (0-8). Blood samples were taken from a group of 10 piglets which did not undergo castration.

**Results** Mean blood cortisol and ACTH concentrations at 30 minutes post-castration were significantly lower in the meloxicam group

than the placebo group ( $p \leq 0.01$ ). The mean haptoglobin concentration at 24 hours was not significantly reduced ( $p = 0.178$ ). The distribution of the GBS during castration was similar in both groups. There were significant differences in the GBS after castration at both 2 and 4 hours post-castration with a greater proportion of piglets in the meloxicam group showing no behavioral alterations (82.7% versus 68.0% at both time points). The score distribution



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J. Dairy Sci. 93:2459-2467  
doi:10.3168/jds.2009-2813  
© American Dairy Science Association®, 2010.

### The effect of meloxicam on behavior and pain sensitivity of dairy calves following cauterity dehorning with a local anesthetic

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## ABSTRACT

Effects of a single injection of meloxicam on calf behavior, pain sensitivity, and feed and water intakes were examined following dehorning. Sixty Holstein heifer calves were blocked by age and randomly assigned to receive an i.m. injection of meloxicam (0.5 mg/kg) or a placebo. All calves were given a lidocaine caudal nerve block (5 mL per horn). Treatments and nerve blocks were administered 10 min before cauterity dehorning. Continuous sampling of behavior was performed during five 1-h intervals using video recordings, and total daily activity was monitored using an accelerometer. A pain sensitivity test was administered with a pressure algometer, and feed and water intakes were recorded daily. Calves were sham-dehorned 24 h before actual dehorning to establish baseline values, and all variables were assessed at the same times following dehorning and sham dehorning for up to 48 h post-dehorning. Meloxicam-treated calves displayed less ear flicking during the 48 h following dehorning

Dehorning refers to amputation of horns in mature cattle or removal of the horn buds of calves; when conducted before 3 wk of age, it is referred to as disbudding. Dehorning is routine management practice on dairy farms, performed to prevent cattle from injuring each other as well as their human handlers. Even though dehorning is done with the long-term welfare of the animals in mind, it does cause acute pain or distress, as indicated by changes in plasma cortisol (Petrie et al., 1996; Sutherland et al., 2002; Heinrich et al., 2009), heart rate (Grandahl-Nilsson et al., 1999; Stewart et al., 2008), eye temperature (Stewart et al., 2008), electroencephalogram (Gibson et al., 2007), and calf behavior (McMeekan et al., 1999; Sylvester et al., 2004). The degree of physiologic and behavior response varies with the method of dehorning. Amputation dehorning caused a greater cortisol response than dehorning by heat cauterization (Petrie et al., 1996).

Local anesthetics such as lidocaine and bupivacaine have been used for reducing physiologic and behavior changes associated with dehorning surgery (Graf and Seiss, 1999), as local anesthetic dissolved, cortisol release (Heinrich et al., 2009) and behaviors such as ear head shaking increased, suggesting the inflammatory pain for up to 24 h (Faulkner and Miller, 2005).

It has been reported that lidocaine injection can reliably described an increase in cortisol immediately following dehorning without treatment associated with acute amputation pain (Faulkner and Miller, 2005). The absence of a dose response when dehorning was performed with lidocaine blockade presented compelling evidence that the pain response was primarily pain related (Faulkner and Miller, 2005).

Behavioral responses such as ear flicking, head shaking, head rubbing, rearing, and



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# Oportunidades de Desarrollo nuevas formas de administración ...





... muchas gracias.

