Genetic improvement strategies for better fertility in dairy cattle



Improving the reliability of fertility breeding values

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Outline

- -Trends in fertility around the world
 - Genetics (additive and non-additive)
- -Improving the reliability of fertility breeding values
 - Genomics and an information nucleus
 - Multi-trait fertility model
 - How collecting more data can help

Calving interval for 15 countries (phenotypic trend)

Dairy Futures





Woolaston et al. (2012)

Calving interval for 15 countries (weighted phenotypic trend)





Genetic Trend for Holstein cows by year of birth



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Summary of worldwide snapshot of fertility

- Evidence for phenotypic trend in calving interval starting to improve
- -Variable evidence for genetic trend in females born
 - Some countries seeing an improvement in the trend between 2000-2005
 - Not consistent in all countries





Other genetic effects

- -Non-additive (or inbreeding and crossbreeding)
- Lethal mutations: fertility "haplotypes" associated with lower (3.1% to 3.7%) conception rates and lower (1.1% to 3.7%) non-return rates



Heterosis effects (calving rate)

Cross	Effect (%)
NZ Holstein x Jersey	+2.2%
N. American Holstein x Jersey	+2.9%
NZ x N. American Holstein	+0.9%

(Pryce et al., 2007: Proc NZSAP)

Reduction in fertility of progeny of non-inbred half-sibs (12.5% inbreeding)



Study	Country	Measure	Effect
Hudson and Van Vleck (1984)	USA	Calving interval	+1.2 d
McParland et al (2007)	Ireland	Calving interval	+8.8 d
Gonzalez-Recio et al. (2007)	Spain	Pregnancy rate	-1.7%*
Hoeschele (1991)	USA	Days open	+1.5 d
Wall et al (2005)	UK	Calving interval	+2.9 d
Pryce et al (2013)	Australia	Calving interval	+2.5 d

* For cows with F between 6.25 and 12.5%

Rate of inbreeding



Study	Country	Breed	Rate of inbreeding/year
McParland et al (2007)	Ireland	Holstein	+0.10%
Wiggans et al (1995)	USA	Holstein	+0.20%
Wiggans et al (1995)	USA	Jersey	+0.22%
Haile-Mariam et al	Australia	Holstein	+0.17%
Haile-Mariam et al	Australia	Jersey	+0.25%

Contribution of inbreeding to phenotypic trend





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10 sires 10 sires

Australian strategy to genetically improve fertility



- Australian Profit Ranking index
- Genomics and an information nucleus
- Multi-trait fertility model
- How collecting more data can help

Development of Australian fertility BVs





Australian Profit Ranking (APR)



Profit-based selection index

- Efficient production
- Survival
- Milking speed
- Temperament
- Fertility
- Cell Count

 Farmers use the APR when selecting bulls





Direction of APR



- Industry criticism that there is "insufficient" weight on fertility in the APR index
- A new approach to selection index.
- What's the cost?



Annual response to selection (unit of measure)



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	Economic Index	APR	
Milk Yield (kg)	62.00	53.52	
Protein Yield (kg)	2.30	2.02	Di
Fat Yield (kg)	2.72	2.40	Ec
Survival (longevity)	1.07	1.10	W
Fertility (%)	0.25	0.51	
SCC (Mastitis Resistance)	2 31	2.86	
Liveweight (kg)	1.00	1.20	
Milking Spood	0.16	0.17	-20.00 -13.00
	0.10	0.17	
remperament	0.37	0.35	

Difference between Economic index and APR worth \$0.83 (<4%)



Improving reliability



Development of Fertility ABVs

2003 Fertility ABV Developed – Calving interval 2010 Increased emphasis on fertility in APR

2003 Fertility Added to APR 2011 Introduction of genomics. Genomic data improves reliability of Fertility ABVs for young bulls

Impact of genomics on reliability of fertility EBVs (young bulls with no progeny)



Country	Pre Genomics	Post Genomics	Change
Australia	16	38	22
Canada	33	60	27
D/S/F			28
Germany	25	43	18
Italy			38
UK			29
USA	31	72	41









3571 bulls with Australian daughters Australian national DNA reference set



8716 cows deliberately selected





Ginfo



Genomic information nucleus

- 10,000 Holstein genomes and Jernomics captured 1 time-point, *Ginfo* is designed to be on-going
- Work with herds with great data, rather than cows with great data
- The reference population needs updating, predictions of genomic breeding values deteriorate as the reference population differentiates from the general population
- Focus on fertility!

Genomic information nucleus



Recruiting of herds started

Opportunity: Focussing on herds with a good track record of recording – an opportunity to broaden breeding values to difficult traits e.g. complex disease traits Genomic information nucleus (n~100)

Herd-testing herds (n~3300)

Non herdtesting herds (n~4000)



• 50 herds in phase 1 Individual farm visits Contact Tail hair collection (farmer or HAA) Lab extract DNA DNA • Farmers continue to collect high quality phenotypes Type and additional phenotypes collected Phenotypes • Better genomic breeding values ABVg

Ginfo process



Development of Fertility ABVs



Multi-trait fertility model





Bulls born since 2000: Fertility breeding values



Breed	Number	Reliability OLD	Reliability NEW	Change
Holstein	2421	61.9	68.3	6.5
Aussie Red	29	62.1	70.8	8.7
Jersey	498	62.4	70.0	7.6



Increasing the reliability of fertility breeding values





Percentage of herds that have insemination data that qualifies for fertility breeding values



DairyFutures CRC 5 1.000.000 800.000 600.000 400.000 200.000 0 Aug 2013

Data, data, data!









From farm.....

To Data Processing Centre



Example herd, data reaching ADHIS before and after our intervention









- Project underway to increase the amount of fertility data that is captured
- Increase of 18% cows with fertility data that qualifies for ABV calculation in the same period for data extracted in August 2012 and March 2013
 - Increased awareness milk recording companies
 - Actively going out and getting the data





Other opportunities



Challenges

- 50K does not capture all genetic variation
- Poor persistence of reliability across generations
- Poor prediction of diverse pedigrees
- Cannot use across breed information

Solution: Whole genome sequence?

Genetic variation captured

Current technology (50K) only captures 60% to 90% of genetic variation (fertility, production)

Whole genome sequence information



- Sequencing still more expensive than SNP chip genotyping
- Alternative strategy
 - Sequence key ancestors, impute genotypes from sequenced animals into all animals genotyped with SNP chips for genomic prediction





1000 bull genomes project

- 16 international partners
- Run 3.0: 427 bulls, 2 cow sequences (15 breeds)
 - 29.1 million SNP
- Early discovery: embryonic lethal recessive mutation
 - INRA collaborators
 - *smc2* gene controls chromosome separation during cell division
 - Mutation phenylalanine -> serine
 - Avoid carrier matings



Name	Fold coverage
Starlite	12.8
Shotime	11.9
Goldsmith	11.8
Gravita	15
Orana	9.5
Beau	12
OVGM	12.3
Goldwyn	22.7
Starbuck	30.3
Rameses	12.4
Donor	15.4
Donante	17.1
Mountain	18.9
Enhancer	16.8
Yukon	19
Gibbon	17
Jocko	15.1
Oman	14.7
Manhattan	17.9
Fatal	16.9
Cash	16.8
Boudewijn	18.5
Sabastian	26.2
Vickai	15.2

Whole genome sequence information



1000 Bull Genomes Project

29.1 million variants

SNP selection

- Biological information
- Very large data sets
- Pathway information

100,000 variants ??

Implementation

 Genomics Evaluations (ADHIS)





Summary

World phenotypic fertility trends look promising World genetic trends are variable In Australia:

- Genomics making a difference to fertility
 - A new genomic information nucleus (Ginfo) established
- A new multi-trait fertility ABV has just been released
- We are working towards getting more mating and pregnancy test data to get the most out of this model!
- Sequencing provides exciting opportunities to improve fertility



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Comparison of European production systems to Australian



Europe

- Range of systems: Total mixed rations, grass
- -Calving all year round (some seasonal: IRL)
- -Welfare and sustainable productions systems
- In most countries milk not as important for GDP
- -Subsidies....

Australia and NZ

- Pasture (NZ), pasture and TMR in Australia
- -Seasonal calving (Spring/Autumn)
- Feed efficiency and productivity
- Milk is a major component of national GDP
- -World milk price

World Holstein Friesian Federation





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World Holstein Friesian Federation



Phenotypic assocation between change in milk volume and calving interval (by country)





\$ Annual response to selection (contribution to profit)



	Economic Index	APR	
Milk Yield	-\$3.27	-\$2.82	
Protein Yield	\$13.80	\$12.09	
Fat Yield	\$4.04	\$3.57	
Survival	\$7.56	\$7.75	
Fertility	\$0.76	\$1.53	
SCS (Mastitis Resistance)	\$0.61	\$0.75	
Liveweight	-\$0.85	-\$1.02	
Milking Speed	\$0.29	\$0.30	
Temperament	\$0.99	\$0.94	
TOTAL	\$23.92	\$23.09	

* Difference between optimal vs APR = \$0.83