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Tools For Managing Genetic Variation

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Genetic Variation

- Genetic variation is a valuable resource
 - » allows further progress
 - » acts as a buffer against unforeseen events
- Requires to be 'treasured'
 - » conserved when not of immediate use
 - » use for generating sustainable genetic gain
 - » not to be wasted



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Genetic Variation

- Genetic variation is a valuable resource
 - » allows further progress
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- Requires to be 'treasured'
 - » conserved when not of immediate use
 - » use for generating sustainable genetic gain
 - » not to be wasted
- But what tools do we need?



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Genetic Variation Within Breed

- Risks from loss of diversity can be measured by 'rate of inbreeding'
 - » denoted ΔF
- Tools for managing genetic variation within a breed manage ΔF



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Why Do We Need New Tools?



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Nicholas & Smith (1987)

- A landmark paper in livestock breeding
- Looked at use of technology in dairy cattle
 - » Multiple Ovulation & Embryo Transfer
 - » Cloning
- Showed increased ΔG through
 - » increased selection intensity
 - » use of relatives for evaluation
- Loss of diversity, ΔF , measured by Wright's formula
 - » classic formula, for random selection!



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Nicholas & Smith (1987)

- This was not unusual pre 1990's
- Theory could predict ΔG
- Theory could not predict ΔF for such schemes!



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What Has Changed?



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New Tools: Theory

- New theories relate both ΔG and ΔF to ‘long-term genetic contribution’
- *An individual’s long term genetic contribution, r_i , is the proportion of the future gene pool contributed by the ancestor*



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New Tools: Theory

$$\Delta G = \sum_{\text{individuals}} r_i a_i$$

$$\Delta F = 1/4 \sum_{\text{individuals}} r_i^2$$

- a_i is the individual's Mendelian sampling term - *it is the bit of variation that makes us unique, different from our full sibs*



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New Tools: Theory

- These advances allow us to plan ahead
 - » predict what ΔF will be in a breeding scheme
 - » predict the consequence of a change of management
 - » predict the consequence of introducing technology



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New Tools: Practice

- But what about breeding decisions day-to-day?
 - » e.g.
 - Which animal should I breed from?
 - Should I use the animal a lot or just a little?
 - This animal is high merit but it already has a full sib selected!



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New Tools: Practice

- Optimum Contribution Selection
- Solves precisely this practical problem
maximise ΔG with ΔF equal to a pre-set value
- Identifies which animals to breed from and the optimum numbers of offspring



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How Does Optimum Contributions Work?

Insight comes from a different Nordic problem!



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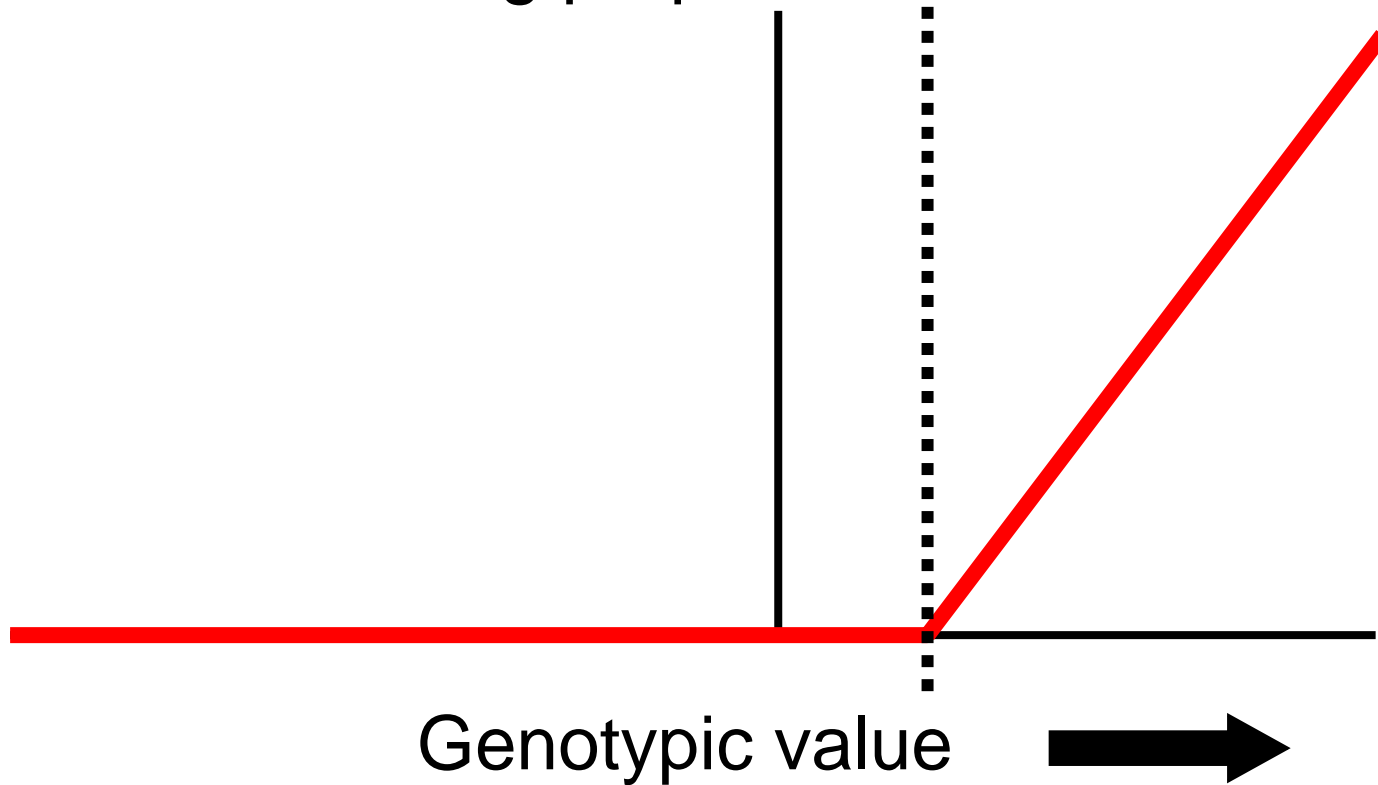
Lindgren's Forestry Problem

- T clones of a tree species
- Each clone has well established genetic value G_i
- Plant a forest with clone i having proportion p_i
- Maximize commercial value $V = \sum_{clones} p_i G_i$
- Minimum diversity required to reduce risks defined by $\gamma < (\sum_{clones} p_i^2)^{-1}$



Bondesson's Solution

Planting proportion

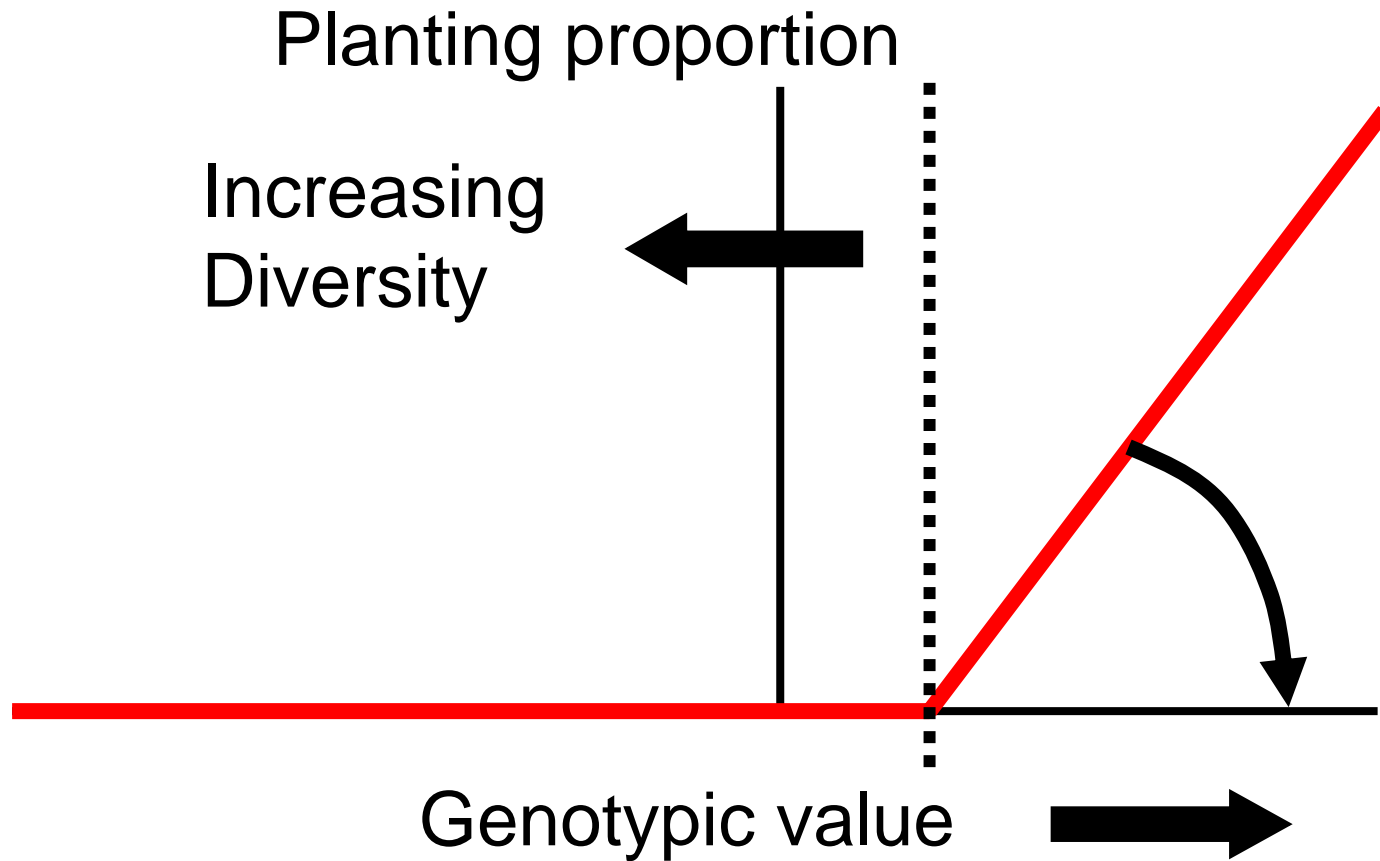


Genotypic value





Bondesson's Solution





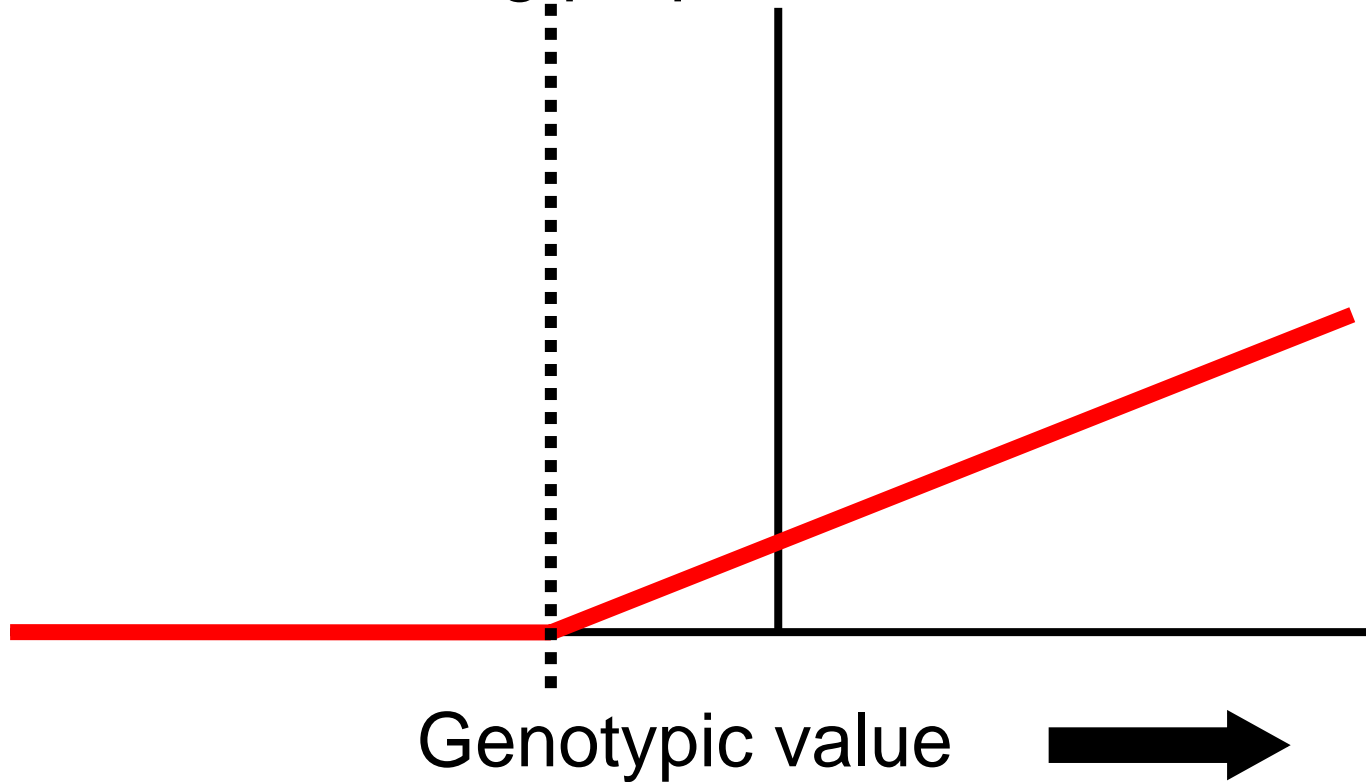
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Bondesson's Solution

Planting proportion



Erling's Celebration

Selection Analogy

- Compare expressions of gain

$$V = \sum_{clones} p_i G_i \quad \& \quad \Delta G = \sum_{individuals} r_i a_i$$

- Compare expressions of diversity

$$\gamma = (\sum_{clones} p_i^2)^{-1} \quad \& \quad \Delta F = 1/4 \sum_{individuals} r_i^2$$

p_i	corresponds to	r_i
G_i	corresponds to	a_i
γ	corresponds to	$(4 \Delta F)^{-1}$



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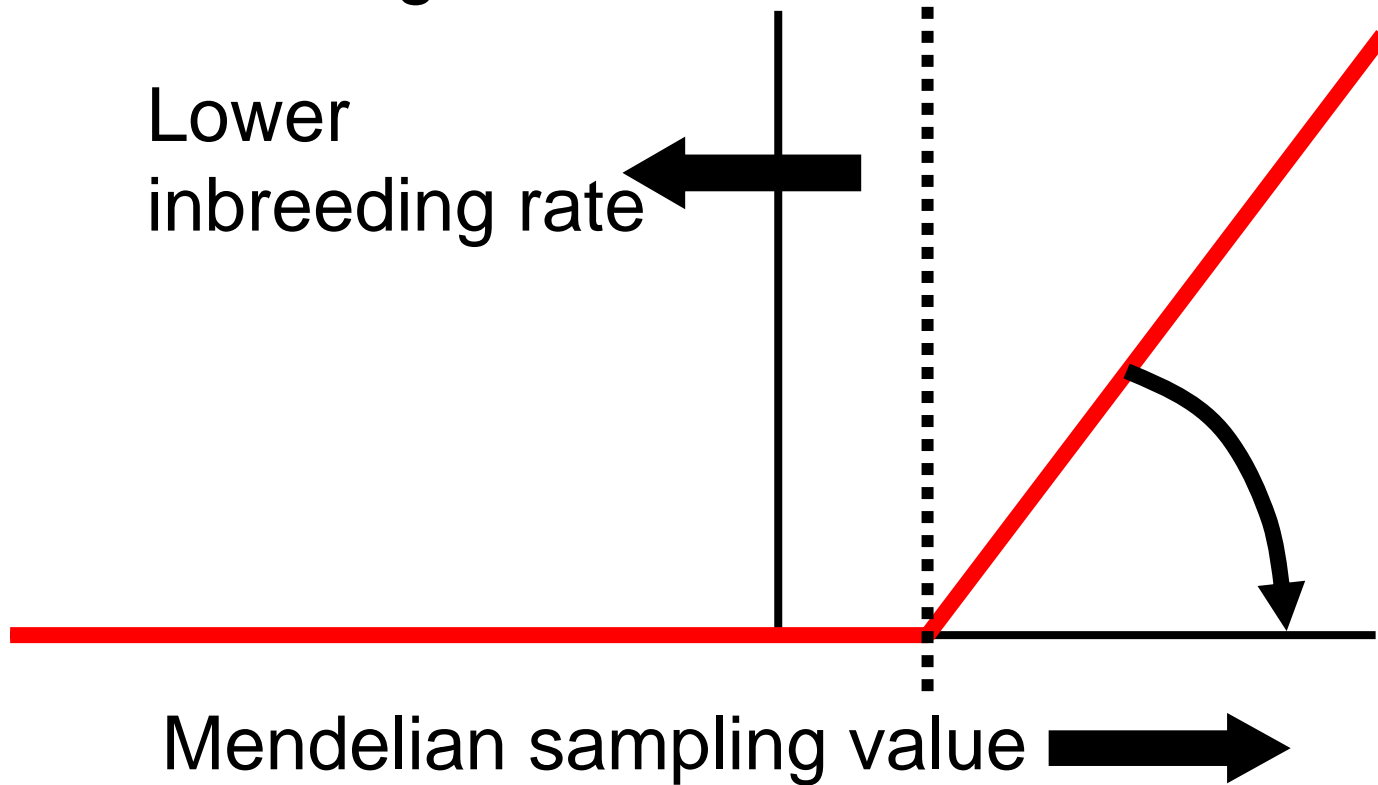


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Ideal Selection

Long term contribution

Lower
inbreeding rate





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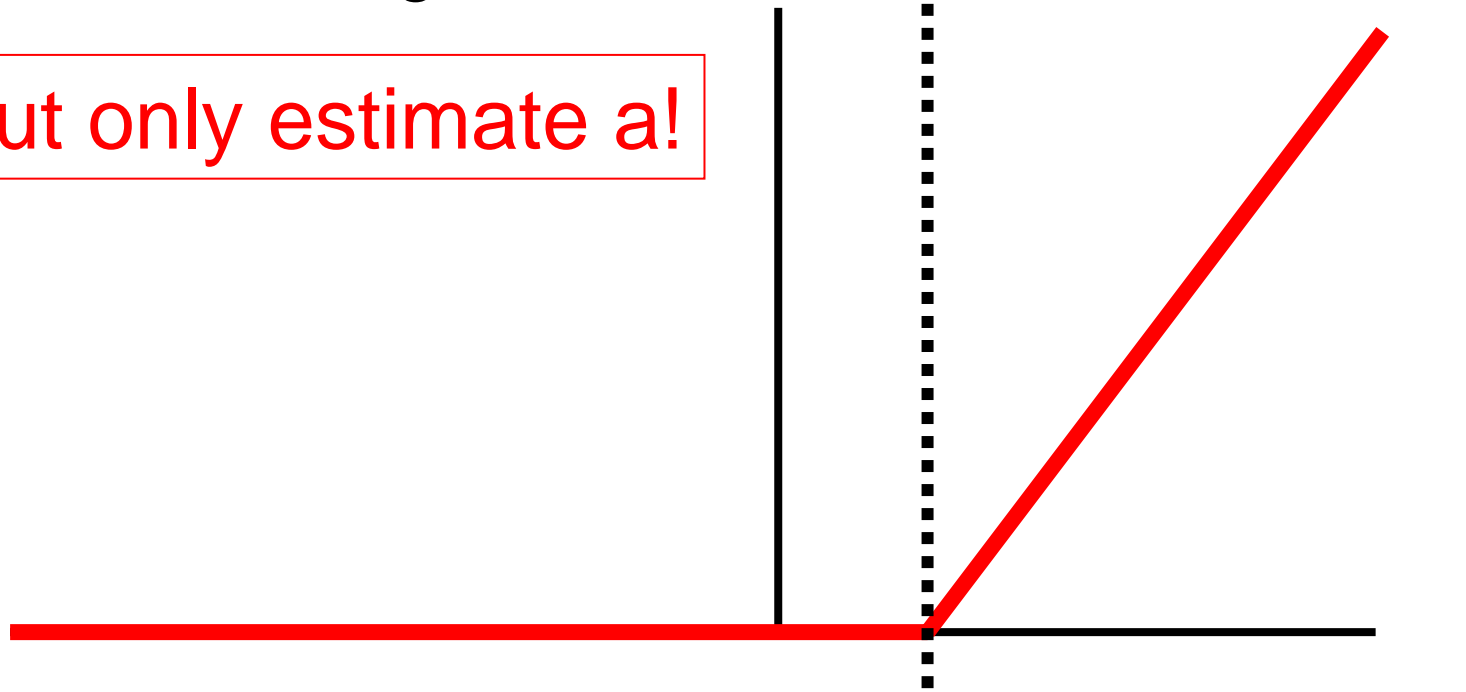


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Practical Management

Long term contribution

But only estimate a!

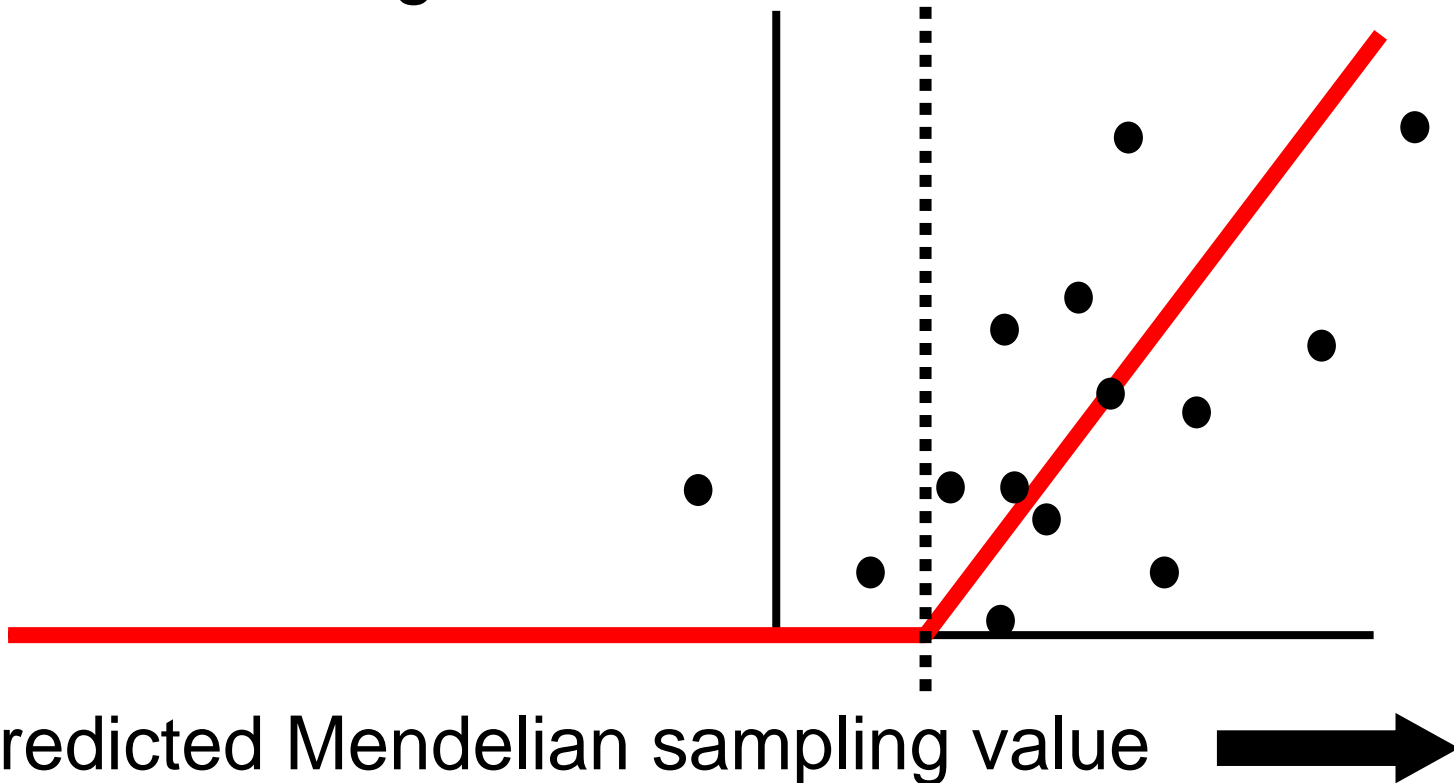


Predicted Mendelian sampling value 



Practical Management

Long term contribution

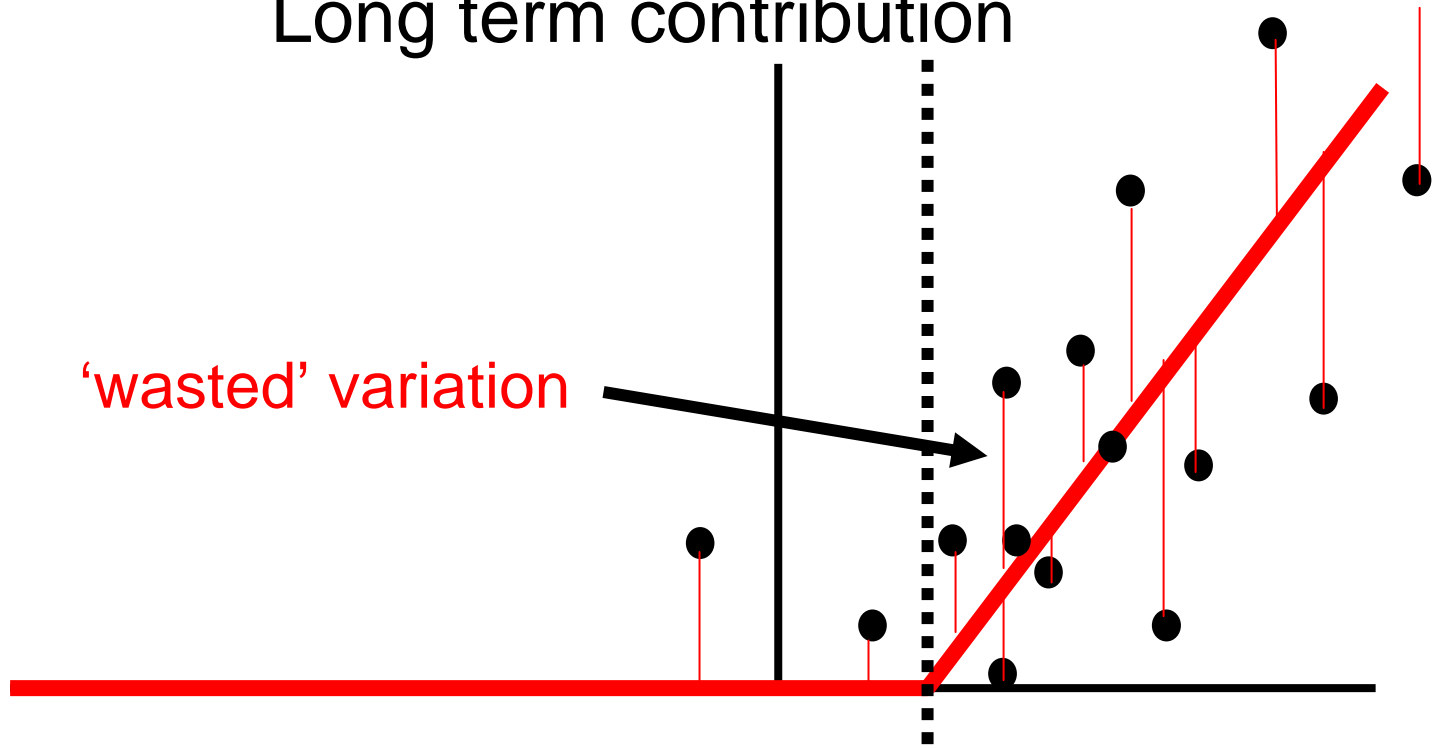




Practical Management

Long term contribution

'wasted' variation



Predicted Mendelian sampling value





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New Tools: Conclusions (1)

- We can manage genetic variation optimally as we make selections
- Historical 'balance' of ΔG with ΔF was seen as losing gain for lower inbreeding
- Optimum Contributions is 'win-win'
 ΔG is maximised for the target ΔF



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New Tools: Conclusions (2)

- There has been another genetic revolution – nothing to do with DNA!
- 20 years ago the chances of selection were determined by the candidates family
 - » the aristocracy
- *Today the chances of selection are determined by what new merit a candidate brings to population*
 - » a meritocracy



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New Tools: Are All Problems Solved?

- 20 years ago could predict ΔG but not predict ΔF
- Now if a breeder uses optimum contributions we know ΔF but cannot predict ΔG !
 - » we know it is best but not how much better



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What New Tools Might The Future Bring?



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The Other Genetic Revolution

- Can genetic selection waste less variation?



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The Other Genetic Revolution

- Can genetic selection waste less variation?
- Yes!
- The DNA revolution has brought with it high throughput cost-effective genotyping
 - » 50,000 marker genotypes per animal for \$200!
 - » rapid results
- Use these for 'Genomic Selection'
 - » use phenotypic data to estimate marker effects
 - » use the estimated marker effects to predict breeding values



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The Other Genetic Revolution

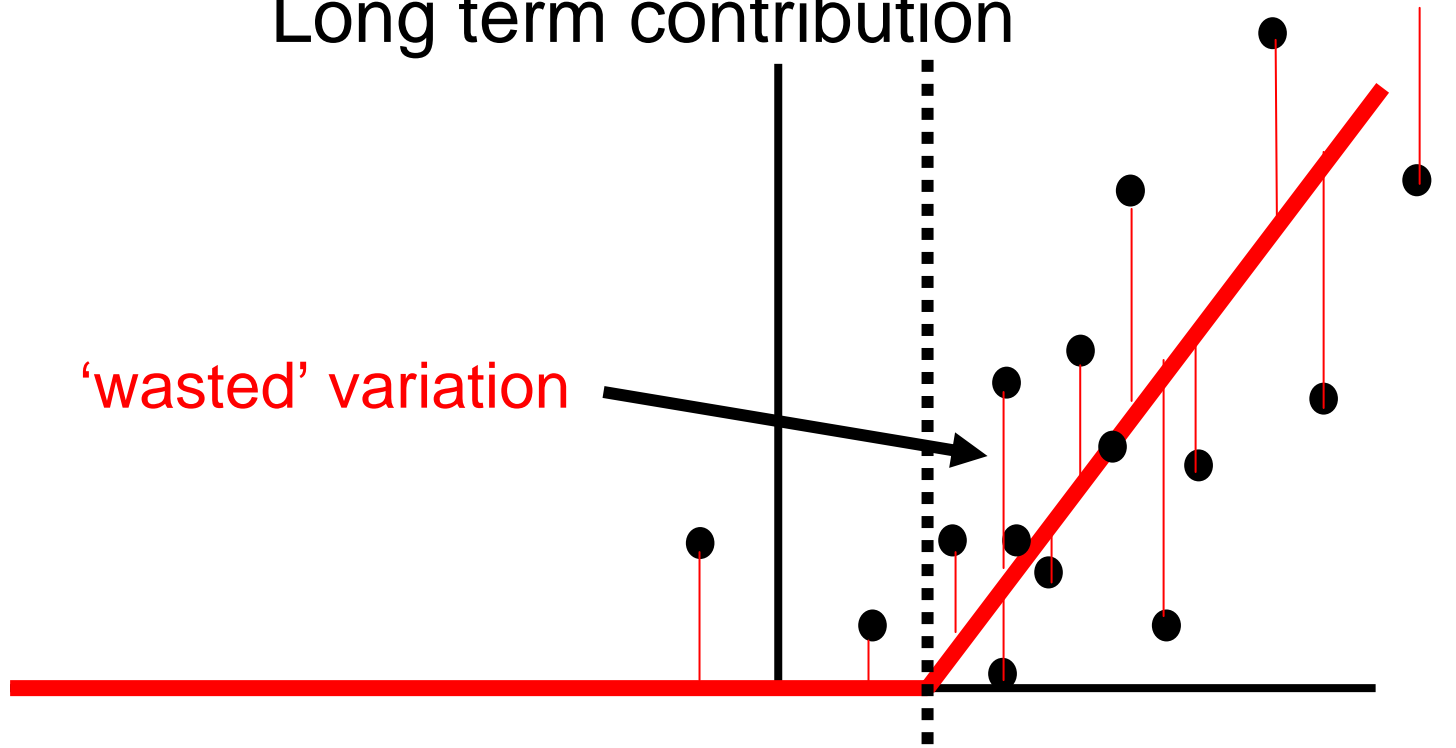
- Genomic Selection produces more accurate breeding values at younger ages
- BUT the additional comes from more accurate predictions of Mendelian sampling values



Without Genomic Selection

Long term contribution

'wasted' variation



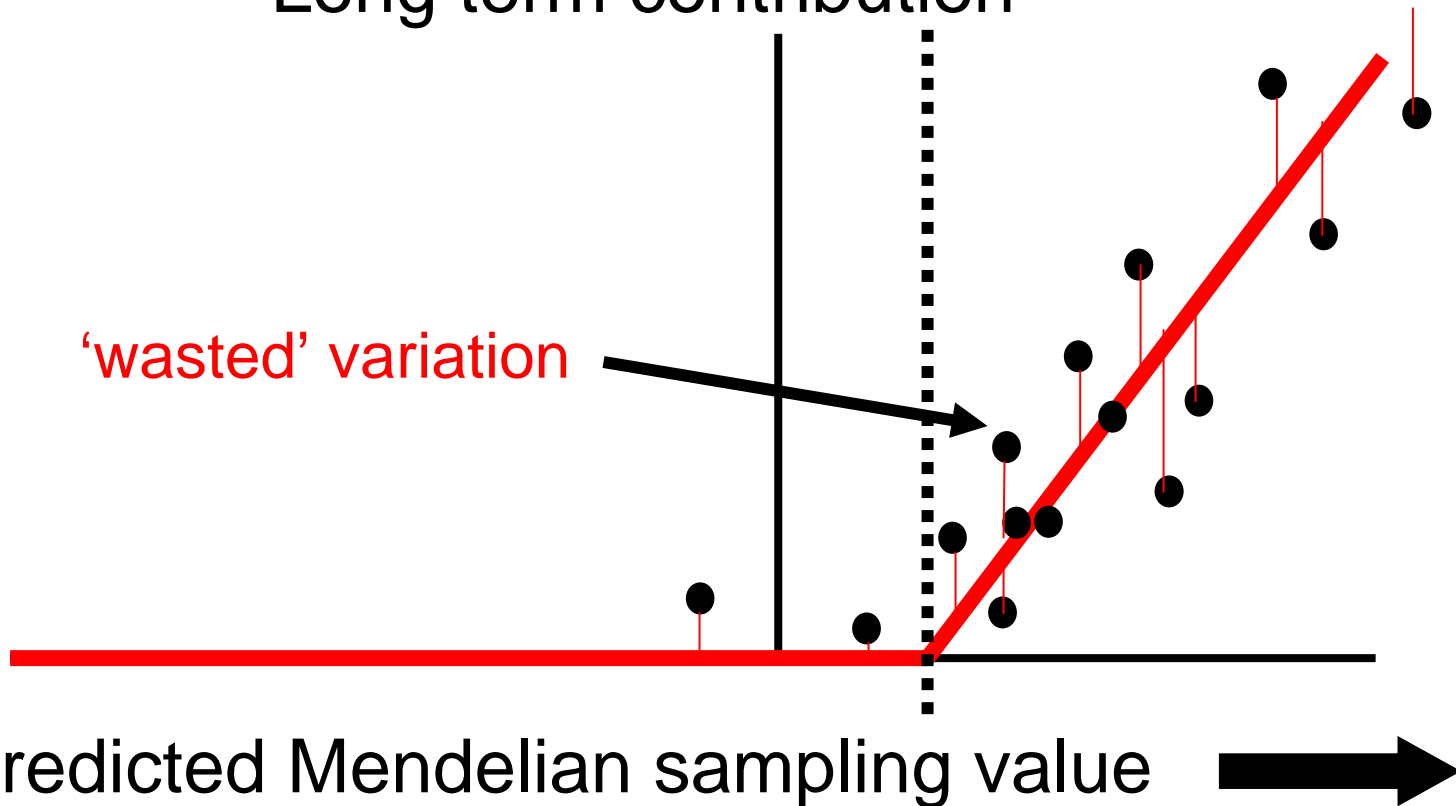
Predicted Mendelian sampling value





Potential With Genomic Selection

Long term contribution





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Potential of Genomic Selection

- Deliver ΔG with less 'wasted variation', less ΔF



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Potential of Genomic Selection

- Deliver ΔG with less 'wasted variation', less ΔF



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Potential of Genomic Selection

- Deliver ΔG with less 'wasted variation', less ΔF
- The challenge of the next decade!



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... And Finally

Erling's Celebration



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Retirement

In my Norsk ordbok

English '*retirement*' = Norsk '*isolasjon*'



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Retirement

In my Norsk ordbok

English '*retirement*' = Norsk '*isolasjon*'

'*isolasjon*' sounds like '*isolation*' in English ???!

I hope I have found the wrong word, Erling!



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Lykke til med fratredelsen din Erling!