SNP based parentage verification via constraint non-linear optimisation

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Interbull meeting, 24-28/10/2016, Puerto Varas, Chile

R. Banks (AGBU)

• "historic" parentage verification (PV)

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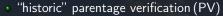
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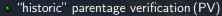
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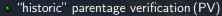
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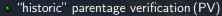
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 - use SNPs for parentage verification directly

SNP parentage verification \rightarrow not too easy

• bi-allelic marker

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 - parents must be among the putative parents \rightarrow cannot be assured
 - likelihood based method adapted from STRs[Boichard et al., 2014]
 - difficult to reject putative parents when N_{SNP} <100

General idea

resort to the linear model

animals marker $= 0.5 \times mum's marker + 0.5 \times dad's marker + mendelian sampling$

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animals marker $= 0.5 \times mum$'s marker+ $0.5 \times dad$'s marker+ mendelian sampling

more mathematical: y = Xb + e

- y : genotype of an animal with one/both parents unknown
- X : matrix of column vectors of genotypes from putative parents
- b : vector of regression coefficients
- e : non-explainable residual

Simple genomic regression

Solve:

 $\arg \min_b f(b) = y'y - 2y'Xb + b'X'Xb \rightarrow \text{simple least square}$

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Simple genomic regression

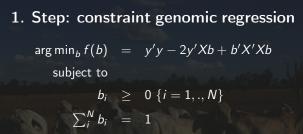
Solve:

 $\arg \min_b f(b) = y'y - 2y'Xb + b'X'Xb \rightarrow \text{simple least square}$

Too simple to be useful:

- b's can have any value ightarrow hard/impossible to interpret
- every b_i can be non-zero \rightarrow who is the parent?

Make it work



Make it work

1. Step: constraint genomic regression arg min_b f(b) = y'y - 2y'Xb + b'X'Xbsubject to $b_i \ge 0 \{i = 1, .., N\}$ $\sum_{i}^{N} b_i = 1$

2. Step: augment X

o column vectors of genotypes of putative parents

population allele frequency vector



Make it work

How to solve:

• 2-step iterative non-linear optimisation solver (NLOpt library[Johnson, 2014])

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Results evaluation

b's assign parentage if $> \frac{1}{3} \rightarrow$ assures number of parents ≤ 2

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How to solve:

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Results evaluation

- *b*'s assign parentage if $> \frac{1}{3} \rightarrow$ assures number of parents ≤ 2
- if both true parents are absent $\rightarrow b$ for population allele frequency vector approaches 1

Test data set

SNP data

• 4612 genotypes of Australian Angus beef cattle

SE!

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 4612 genotypes of Australian Angus beef cattle
 47702 SNPs extracted from Illumina 50K Bead Chip genotypes

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SNP data

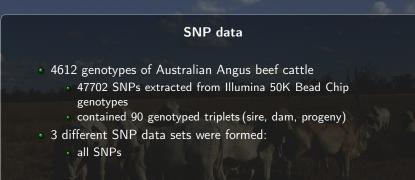
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 - contained 90 genotyped triplets (sire, dam, progeny)

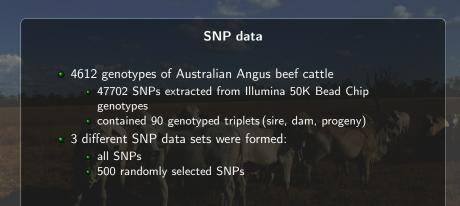
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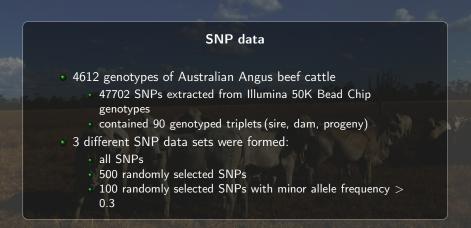


4612 genotypes of Australian Angus beef cattle 47702 SNPs extracted from Illumina 50K Bead Chip genotypes

- contained 90 genotyped triplets (sire, dam, progeny)
- 3 different SNP data sets were formed:







Test data set

Equation data

- $y \rightarrow$ progeny from triplet
- $X \rightarrow$ three different types per y

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dam

- 10 randomly selected animals
- population allele frequency

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X_R 12 randomly selected animals

 population allele frequency

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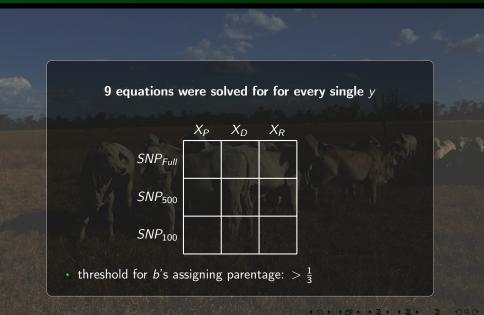
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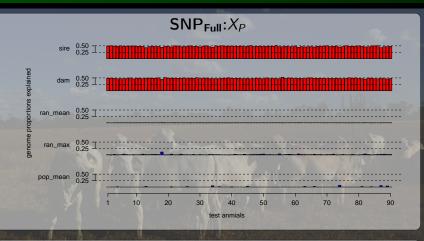
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 - selected animals
- population allele frequency

- randomly selected animals: excluded parents, full sibs and half sibs; re-sampled for every y
- population allele frequency vector was calculated excluding genotypes in X and y

Trial summary



Results



• all parents correctly identified

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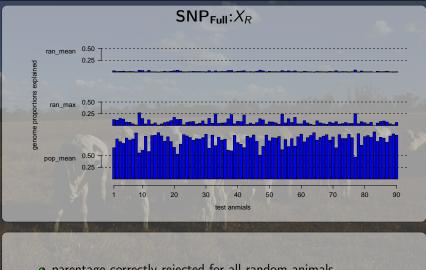
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- all dams correctly identified
- · parentage correctly rejected for all random animals

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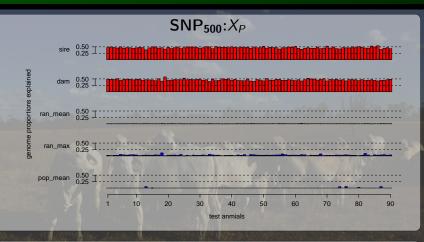
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SNP based parentage verification

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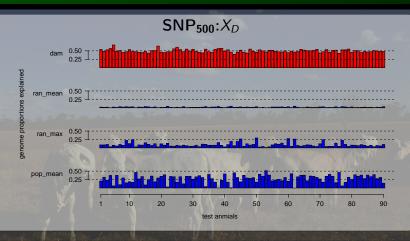


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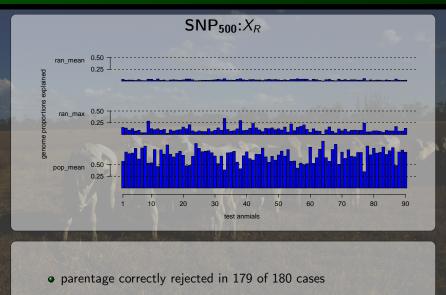
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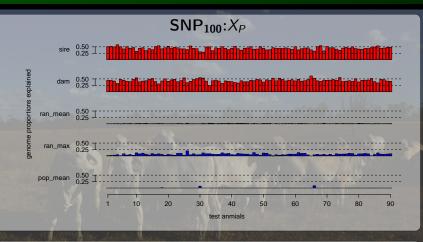


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SNP based parentage verification

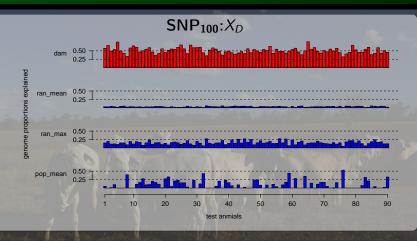
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Results



- parents correctly identified in 173 of 180 cases
- · parentage correctly rejected for all random animals

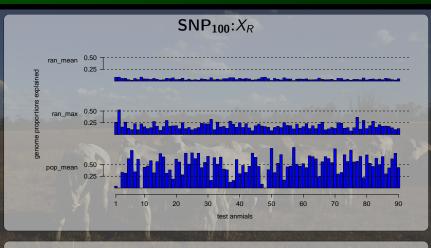
Results



- dams correctly identified in 89 of 90 cases
- · parentage correctly rejected for all random animals

SNP based parentage verification

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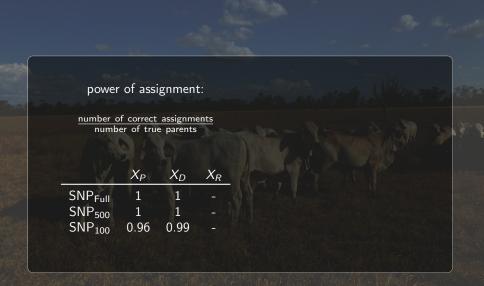


• parentage correctly rejected in 178 of 180 cases

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SNP based parentage verification

Results summary



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power of assignment:				power of exclusion:				
	number of correct assignments number of true parents				$1-rac{number\ of\ wrong\ assignments}{number\ of\ possible\ parents}$			
	X _P	X _D	X_R		X_P	X_D	X_R	
SNP _{Full}	Х _Р 1	$\frac{X_D}{1}$	X _R	SNP _{Full}	$\frac{X_P}{1}$	$\frac{X_D}{1}$	$\frac{X_R}{1}$	
SNP _{Full} SNP ₅₀₀		1.12	X _R - -	SNP _{Full} SNP ₅₀₀	1	199302-36	Contractor and the second	

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Discussion

• general performance

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- general performance
 - almost perfect assignment for $N_{SNPs} \ge 500$

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 - may also allow to reject "tricky" sets of putative parents containing progeny, full sibs or half sibs

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