Genome-Wide Selection

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evolution of QTL models

original ideas focused on rare & costly markers models & methods refined as technology advanced

- single marker regression
- QTL (quantitative trait loci)
 - single locus models: interval mapping for QTL
 - QTL model search: QTLs & epistasis
- polygenes (association mapping)
 - adjust for population structure
 - capture "missing heritability"
- \cdot genome-wide selection

what is genomic selection?

use statistical modeling to predict how a plant will perform before it is field-tested

- genomic selection (GS)
 - marker assisted selection (MAS)
 - genome-wide selection (GS)
- other uses of word (relevent to systems genetics)
 - natural selection: survival of the fittest
 - model selection: search for QTLs
 - selection bias: overestimate of QTL effects

why use genomic selection?

- trait is highly polygenic (genetically variable)
 - influenced by a few key genomic regions
 - high heritability (low environmental variation)
- measuring trait is costly
 - difficult or expensive process (technology)
 - measuring tool may be highly variable
 - time-consuming (plant has to grow first)
 - desire to streamline multi-year selection

what is genomic selection?

- forms of genome-wide selection
 - marker-assisted: with phenotypes
 - marker-based: without phenotypes
- use markers to improve selection for complex traits
 - predict phenotype from marker genotype
 - select candidates based on best marker genotypes
 - use training set to predict test set of individuals

old paradigm: marker prediction

- 1990s & 2000s: markers were expensive
- economic strategy:
 - first identify significant markers (QTL analysis)
 - use best markers to genotype selection candidates
- estimate marker effects by multiple regression
 - treat genetic effects as fixed and few
 - $E(y) = \mu_q, q = (q_1, q_2, q_3)$

marker assisted selection (MAS)

EARLY GENERATION SELECTION

PEDIGREE METHOD MARKER ASSISTED SELECTION P1 P2 х P1 P2 х F1 F1 000000000000000 Phenotypic 000000000000000 F2 screening MAS 00000000000000000 F2 00000000000000000 00000000000000000 Plants spaceplanted in rows for Only desirable F3 F3 individual plant lines planted in F3 selection field Families grown in Families grown in F4 F4 progeny rows for progeny rows for selection. selection. Pedigree selection F5 based on local F5 needs × 7 Preliminary yield trials. Select single F6 F6 plants. × × Further yield F7 trials F7 Multi-location testing, licensing, seed increase Multi-location testing, licensing, seed increase F8 – F12 and cultivar release F8 – F12 and cultivar release

www.21stcentech.com/heard-marker-assisted-breeding/

new paradigm: use "all" markers

- new paradigm with technology advances
 - improved statistical methods and software
 - cheap markers
- using only significant markers to predict trait ...
 - gives good estimates (maybe) of markers ...
 - but does not maximize accuracy
- simple but effective approach
 - treat marker effects as random
 - use all markers (away from QTL if any)

old vs new



9/12

mixed model approach

MAS approach $y = \mu_q + e$, $V(e) = \sigma^2 I$

- estimate fixed QTL effects $\hat{\mu}_q$ (MLEs)
- predict phenotype using fixed effects $\hat{y} = \hat{\mu}_q$

GS approach $y = \mu + g + e$, $V(g) = \sigma_g^2 K$

- estimate kinship *K* from all markers *M* as for poly
- predict random effect \hat{g} using BLUP
- predict phenotype $\hat{y} = \hat{g}$

genomic prediction

- DO example
- rrBLUP fit without QTL
- correlation 0.79



poly + QTL genomic prediction

- DO example
- rrBLUP fit with QTL
- correlation 0.74

