

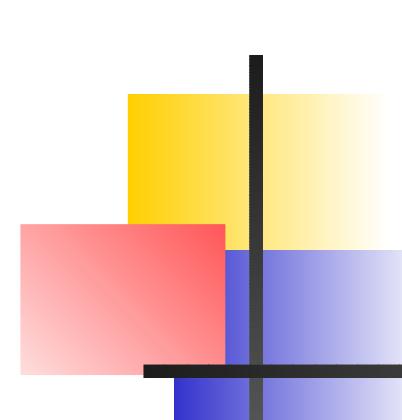
Exercises

- Coopworth data set - see Reference manual
- Five traits with varying amounts of data.
- No depth of pedigree (dams not linked to sires)
- Do univariate analyses
- Do bivariate analyses.
- Use COOP data set and attempt multivariate models.



Multivariate analysis

- Modelling becomes more difficult as number of traits increases - there may be variance but the matrix may not be positive definite (covariances too big).
 - Maybe do bivariate pairs first
 - !GP constrain positive definite - will use EM updates if AI updates generate NPD matrix
 - try FA or CHOL reduced parameterization
 - try Singular XFA or CHOL paramaterization



Usual process

- Univariate analysis of each trait.
 - Identify an appropriate fixed model
 - Check for outliers and problems with data structure
 - Sort out fixed model appropriate for each trait
 - Ensure there is variance at each level: if there is no sire variance in a univariate model, ASReml will not be able to estimate it in a multivariate model

Output:ANOVA

	ANOVA	NumDF	DenDF	F-incr	Pro
9	Trait	2	4.9	5019.79	<.00
10	Tr.line	4	5.3	2.65	0.15
12	at(Tr,2).Age	1	58.5	6.33	0.01

Output:structures

- Covar/Var/Corr UnStructured
132.4 -0.3319
-98.03 659.0
- Covar/Var/Corr UnStructured
27.20 -0.2354
-12.20 98.72

Output:components

Source	Model terms	Gamma	Component	C/SE	% C	U/L
Residual	UnStru 1 1	132.370	132.370	5.29	0	U
Residual	UnStru 2 1	-98.0288	-98.0288	-2.27	0	U
Residual	UnStru 2 2	659.005	659.005	5.19	0	U
Tr.sire	UnStru 1 1	27.2002	27.2002	1.02	0	U
Tr.sire	UnStru 2 1	-12.1986	-12.1986	-0.30	0	U
Tr.sire	UnStru 2 2	98.7165	98.7165	0.88	0	U

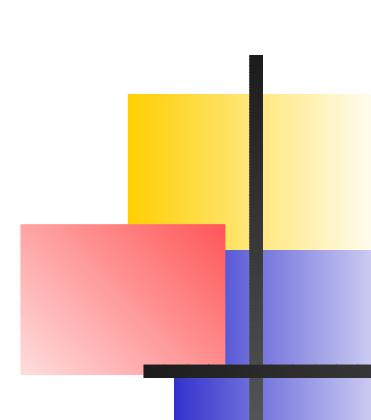
Output:convergence

- 1 LogL=-466.835 S2=1.0000 123 df
- 2 LogL=-451.914 S2=1.0000 123 df
- 3 LogL=-437.150 S2=1.0000 123 df
- 4 LogL=-428.395 S2=1.0000 123 df
- 5 LogL=-427.247 S2=1.0000 123 df
- 6 LogL=-427.201 S2=1.0000 123 df
- 7 LogL=-427.201 S2=1.0000 123 df
- Analysis on Variance scale
 $V = \sigma^2(R + ZGZ')$ with $\sigma^2 = 1$.



Missing data

- Multivariate analysis, with US error variance matrix, ASReml will automatically handle missing values in the traits.



Notes

- R structure is 65×2 because data is ordered traits within records
- G structure is 2×9 because Tr.sire means the effects are ordered sires within traits
- Initial values must be supplied but are often difficult to guess. Specifying them as zeros (3*0) tells ASReml to work out some values from the data.

1 G Structure

- ADG WT ~ Trait Tr.line,
at(Tr,2).Age !r Tr.sire
1 2 1
0 # 65 records
- Trait 0 US
- 3*0
- Tr.sire 2 # =====
- Tr 0 US # =====
- 3*0 # =====
- sire 0 ID # =====

1 R structure

```
■ ADG WT ~ Trait Tr.line,  
           at(Tr,2).Age !r Tr.sire  
   1 2 1  
   0      # 65 records # <=====  
Trait 0 US          # <=====  
3*0                # <=====  
Tr.sire 2  
Tr 0 US  
3*0  
sire 0 ID
```

Variance header line

- ADG WT ~ Trait Tr.line,
at(Tr,2).Age !r Tr.sire
1 2 1 # <=====
0 # 65 records
- Trait 0 US
- 3*0
- Tr.sire 2
- Tr 0 US
- 3*0
- sire 0 ID

Bivariate continued

- Now stack WT effects below ADG effects:
 $\mathbf{u} = \text{vec}(\mathbf{U})$
 $\text{var}(\mathbf{u}) = \Sigma_C \otimes \Sigma_R$
- Essential to get order correct: second term is nested within first.
- Adding part 3 to the Harvey job: the model

! PART 3

ADG WT ~ Trait Tr.line,
at(Tr,2).Age !r Tr.sire

Bivariate Analysis

- With Harvey data, could analyse ADG and WT - to get two sets of sire effects and two sets of residuals.
- Consider sire effects (U) in a 9×2 table.
Rows are independent: $\Sigma_R = I_9$
Columns are correlated but have different variances: $\Sigma_C = \begin{pmatrix} S_{11} & S_{12} \\ S_{21} & S_{22} \end{pmatrix}$



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B4 Bivariate Analysis

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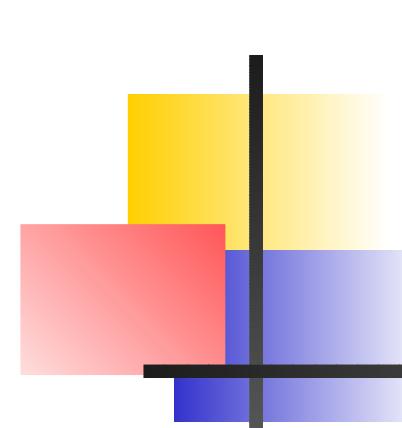


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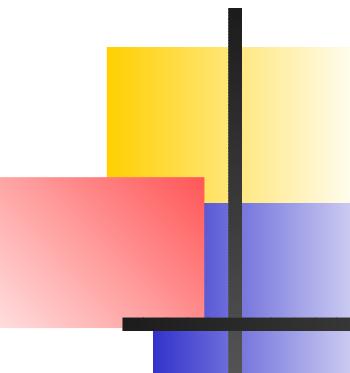
Sire model → Animal model

- Solving the model in terms of genetic components can lead to further problems.
 $\sigma_E^2 = \sigma_e^2 - 3\sigma_s^2$ may be negative and not estimable (ASReml requires a positive residual variance)
- $h^2 = 4\sigma_s^2/(\sigma_e^2 + \sigma_s^2)$ is ≤ 4 but if the genetic model is correct, should be less than 1.
- I.e. Our model may not adequately represent the variation in the data leading to unacceptable genetic parameter estimates.



Estimability of Components

- Hendersons method III equated the Sire Mean Square (SMS) to its expectation $\sigma_e^2 + k\sigma_s^2$
- While $\text{SMS} \geq 0$,
 $\sigma_s^2 = (\text{SMS}-\text{EMS})/k$ is $\geq -\text{EMS}/k$
- In REML, often constrain components to be positive (equivalent to estimating σ_e^2 after dropping sire from the model).



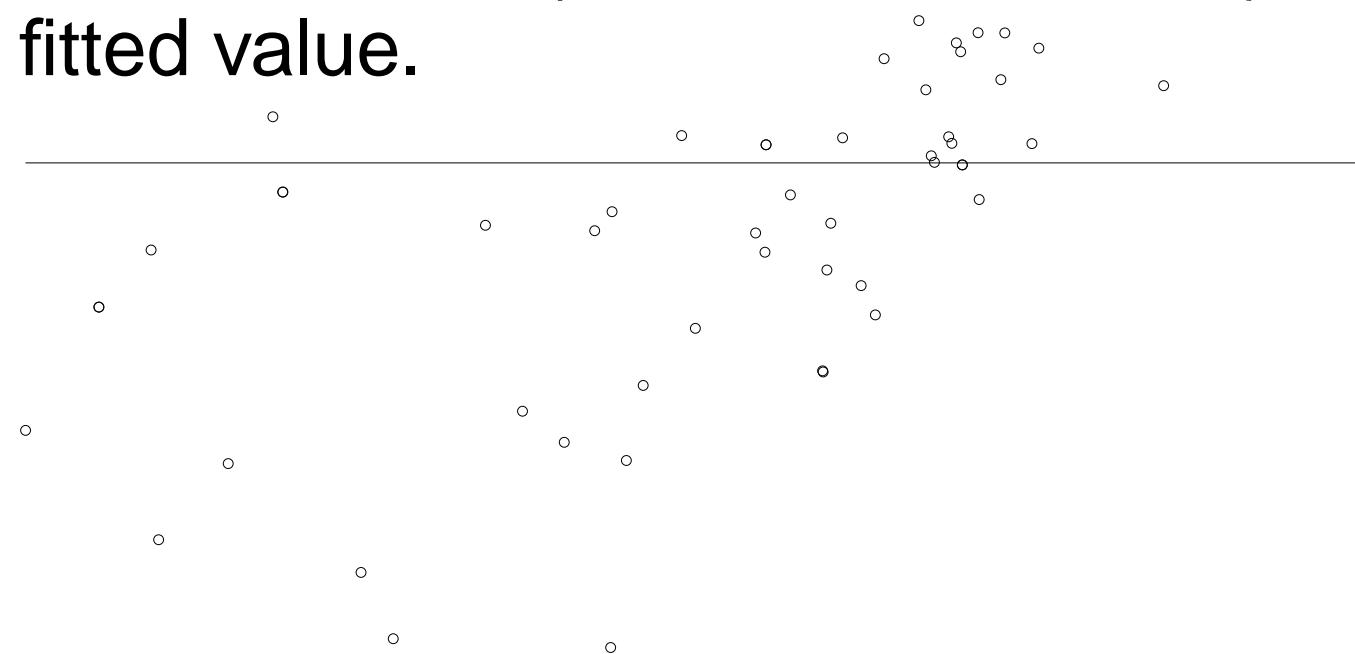
Other genetic components

- Genetic maternal definition `DAM !P`
and include DAM in the model
- Permanent environment effect
`Use ide(animal)`
- Maternal environment effect
`Use ide(DAM)`

Plot of Residuals

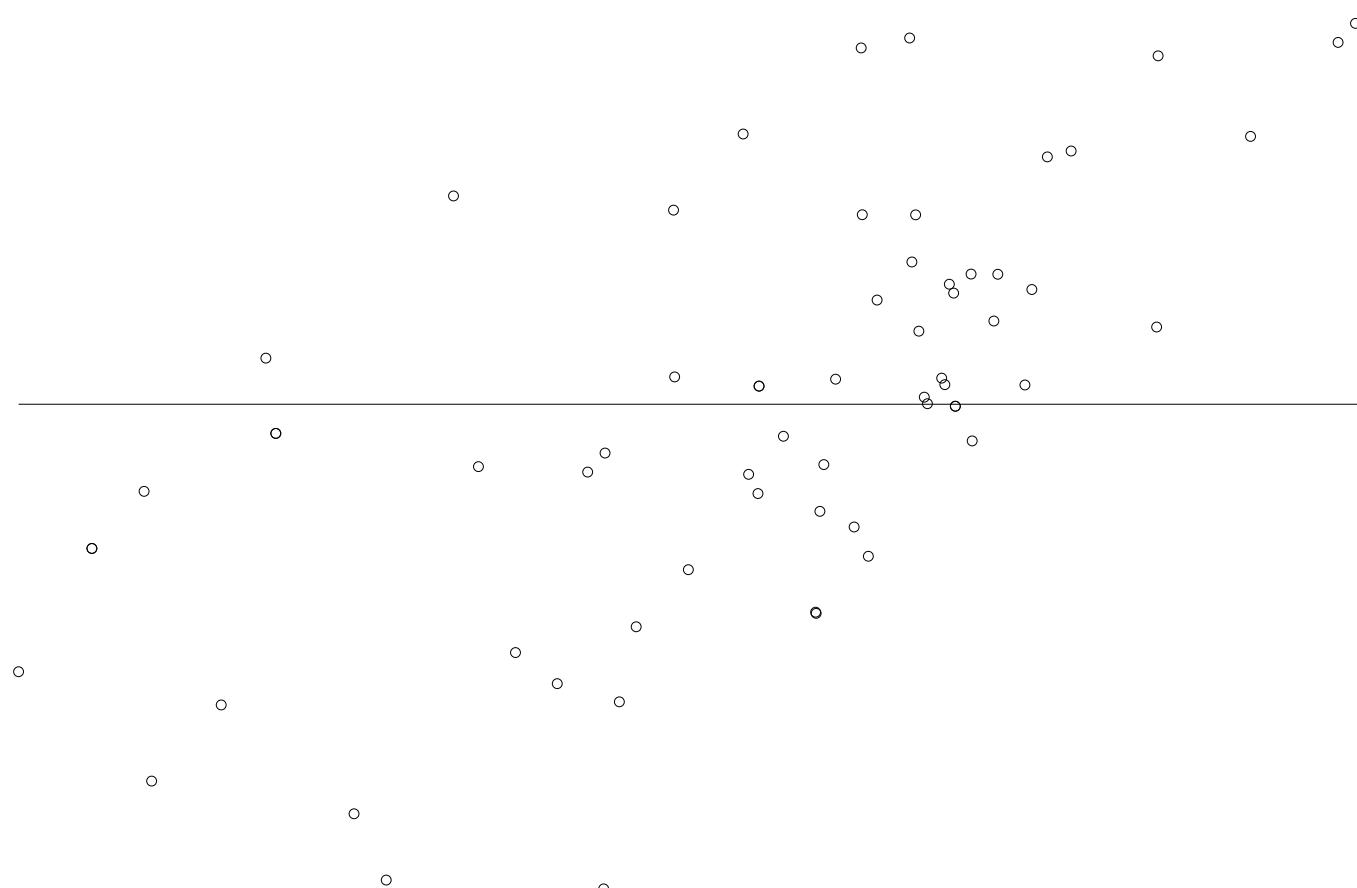
Harvey Test data - Animal Model Residuals vs Fitted values
Residuals (Y) -8.84: 6.95 Fitted values (X) 148.88: 199.43

- Apparent mean variance relationship arises because the animal model adds a proportion of the residual (from the sire model) into the fitted value.



Plot of Residuals

Harvey Test data - Animal Model Residuals vs Fitted values
Residuals (Y) -8.84: 6.95 Fitted values (X) 148.88: 199.43



Results - Effects

			Estimate	StandErr	T-val
5	DamAge	1	-1.47752	1.88080	-0.79
4	line	2	-14.4126	6.28603	-2.29
		3	6.46567	5.29341	1.22
9	mu	1	183.489	9.31871	19.69
1	animal		74	effects fitted	

Results - ANOVA

■ ANOVA	NumDF	DenDF	F-incr	Prob
9 mu	1	5.9	5906.95	<.001
4 line	2	5.9	6.19	0.035
5 DamAge	1	57.8	0.62	0.435

is the same as the Sire model

Results - components

- 5 LogL=-188.781 S2=54.092 61df 1.957
- 6 LogL=-188.777 S2=47.314 61df 2.411
- 7 LogL=-188.777 S2=46.100 61df 2.506
- Final parameter values 2.508

Source	terms	Gamma	Component	C/SE
animal	74 74	2.5086	115.578	1.04
Variance	65 61	1.0000	46.0723	0.51

Running the model in ASReml

```
■ !REDO !ARG !? 1=Sire 2=Animal  
Harvey Test data - Sire and Animal M  
animal !P sire 9 dam line 3 # Added  
DamAge ADG Age WT  
harvey.dat      # Pedigree file line a  
harvey.dat      !DOPART $1 # Data  
!PART 1  
ADG ~ mu line DamAge !r sire  
!PART 2  
ADG ~ mu line DamAge !r animal
```

Variance model

- Fitting an animal model for the harvey data

$$\mathbf{V} = \sigma_E^2 \mathbf{I}_{65} + \sigma_A^2 \mathbf{Z} \mathbf{A} \mathbf{Z}'$$

where $\mathbf{Z} = (\mathbf{0}_{65 \times 9} \ \mathbf{I}_{65})$

Thus \mathbf{V} simplifies to

$$\sigma_E^2 \mathbf{I}_{65} + \sigma_A^2 (0.25 \mathbf{Z}_s \mathbf{Z}'_s + 0.75 \mathbf{I}_{65})$$

$= (\sigma_E^2 + 0.75\sigma_A^2) \mathbf{I}_{65} + 0.25\sigma_A^2 \mathbf{Z}_s \mathbf{Z}'_s$ which is exactly the same as under the sire variance model.

reducing to

- $A_i^{-1} = \begin{pmatrix} A^{-1} + pqp' & pq \\ qp' & q \end{pmatrix}$ where p is zero except for two $\frac{1}{2}$'s in parental rows,
 $q = (1 + f_i - p'Ap)^{-1} = (1 - (a_{s,s} + a_{d,d})/4)^{-1}$ so that it just requires keeping $\text{diag}(A)$.
- For the case of a sire model (dams unknown, sires not inbred),

$$A = \begin{pmatrix} I_9 & 0.5Z'_s \\ 0.5Z_s & 0.25Z_s Z'_s + 0.75I_{65} \end{pmatrix}$$



Relationship Matrix

- The diagonal of the relationship matrix ($a_{i,i}$) is $1 + f$ where $f = a_{s,d}/2$ is the inbreeding coefficient.
- The relationship of an animal with other animals is the average of its parental values.
- When parents are listed before progeny, this gives a straight forward way to calculate relationships.
- However, the inverse relationship matrix is in fact easier to calculate and more sparse.



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B3 Animal model

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Line+Sire

■ predict Sire !present Sire Line

3.0000	173.4734	4.1885 E
4.0000	157.4867	3.6989 E
5.0000	167.7099	3.8875 E
6.0000	183.5371	3.8512 E
7.0000	186.8966	3.4811 E
8.0000	183.8807	3.6774 E
9.0000	179.5918	3.4811 E

Overall Stnd Error of Diff 5.146

Line+Sire

- predict Sire !present Sire Line
Line is average of combinations present
DamAge is evaluated at 4.3846

Sire	Pred_Value	Stand_Error	Ecode
1.0000	176.9570	3.5634	E
2.0000	180.6023	3.5962	E
3.0000	173.4734	4.1885	E
4.0000	157.4867	3.6989	E
5.0000	167.7099	3.8875	E
6.0000	183.5371	3.8512	E

Predict

- predict Line

Predicted values of ADG

DamAge evaluated at average value 4.
Sire is ignored in the prediction

Line	Predicted_Value	Stand_Error	Ec
1.0000	177.0109	4.0193	E
2.0000	162.5983	4.8301	E
3.0000	183.4766	3.4415	E
Overall	Stnd Error of Diff	5.851	

Fixed effects

■ Term	Level	Effect	SE
DamAge	1	-1.478	1.881
Line	1	0.000	0.000
Line	2	-14.41	6.286
Line	3	6.466	5.293
mu	1	183.5	9.319

■ Notes:

Terms are in reverse order (ASReml solves from bottom)

Line-1 effect is singular.

ANOVA

- Degrees of Freedom and Stratum Variances

5.93	341.034	7.2	1.0
55.07	132.756	0.0	1.0

ANOVA	NumDF	DenDF	F-incr	Prob
9 mu	1	5.9	5906.95	<.001
4 Line	2	5.9	6.19	0.035
5 DamAge	1	57.8	0.62	0.435

- Damage is NS; Line is significant tested against Sire variance.

Sire BLUPs

- from the .sln file – sum to zero within lines

Sire	1	-0.5386E-01	4.137
Sire	2	3.591	4.142
Sire	3	-3.538	4.269
Sire	4	-5.112	4.468
Sire	5	5.112	4.468
Sire	6	0.6051E-01	4.055
Sire	7	3.420	3.914
Sire	8	0.4042	3.990
Sire	9	-3.885	3.914

Genetic parameters

■	3	PhenVar	1	161.7	35.07
	4	GenVar	1	115.6	110.8
Heritability = GenVar 4 / PhenVar 3					
= 0.7150 0.5877					

Notice: The parameter estimates are followed by their approximate standard errors

Genetic components

■ Create a PIN file (harvey.pin)

```
#           1 is Sire component  
#           2 is Residual  
F PhenVar 1 2    #3 is Sire + Residual  
F GenVar 1*4.    #4 is Sire x 4.0  
H Herit 4 3      #Heritability is GenVar/Phe
```

■ Run using ASReml -p harvey

This extracts the variance components from the .asr file and their variances from the .vvp file and computes the requested quantities

Components

- 5 LogL=-188.777 S2= 132.76 61 df 0.2176 1.000

Source	terms	Gamma	Component	Comp/SE	% C	
Sire	9	9	0.217651	28.8946	1.04	0 P
Variance	65	61	1.000000	132.756	5.25	0 P

- Notice $\gamma = 0.21765$, $\sigma_e^2 = 132.756$,
 $\sigma_s^2 = \gamma\sigma_e^2 = 28.8946$.

Random design

- Z has 9 columns being zeros except for 8 8 5 8 7 6 8 7 8 1's respectively indicating which sire
- The variance model is $\sigma_e^2(I + \gamma ZZ')$ where $\sigma_s^2 = \gamma\sigma_e^2$
- The genetic model
 $\sigma_A^2 = 4\sigma_s^2; \sigma_s^2 = 0.25\sigma_A^2$
 $\sigma_E^2 = \sigma_e^2 - 3\sigma_s^2; \sigma_e^2 = \sigma_E^2 + 0.75\sigma_A^2$



Fixed Design

- X has 5 columns
 - mu is a column of ones
 - line-1 has 21 1's, 15 0's, 29 0's
 - line-2 has 21 0's, 15 1's, 29 0's
 - line-3 has 21 0's, 15 0's, 29 1's
 - DamAge (covariate) has vector of dam ages
- This design has 1 singularity because the three line columns sum to give the mu column.
- ASReml will set $\tau_2 = 0$

Summary

■ Model term	Size	#mv	#00	MinNon0	Mean	MaxNon0
1 ID	0	0	101.0	133.0	165.0	
2 Sire	9 0	0	1	5.0154		9
3 Dam	0	65	0.000	0.000	0.000	
4 Line	3 0	0	1	2.1231		3
5 DamAge	0	0	3.000	4.385	5.000	
6 ADG Variate	0	0	144.0	176.6	206.0	
7 Age	0	0	337.0	416.8	498.0	
8 InitialWT	0	0	144.0	241.1	300.0	

2.2 Sire model

- Harvey Test data - Sire Model
 - animal sire 9 dam line 3
 - DamAge ADG Age WT
 - harvey.dat
 - ADG ~ mu line DamAge !r sire



Harvey.dat summary

- 9 sires representing 3 sire lines
(1 2 3) (4 5) (6 7 8 9)
8 8 5 8 7 6 8 7 8 records (65)
- Data set originally distributed with Harvey's program

Harvey.dat

■ animal sire dam

	line	Dam	Age	ADG	Age	WT		
101	1	0	1	3	192	390	224	1
102	1	0	1	3	154	403	265	1
103	1	0	1	4	185	432	241	1
104	1	0	1	4	183	457	225	1
105	1	0	1	5	186	483	258	1
106	1	0	1	5	177	469	267	1



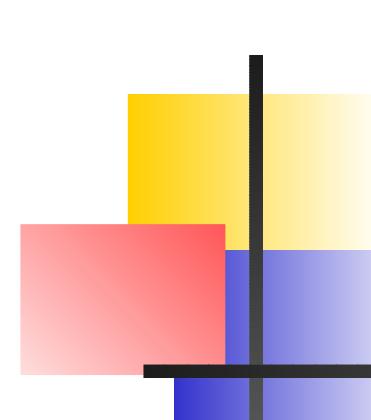
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B2 Sire Model

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The challenge

- to define X Z R and G to obtain the desired analysis.

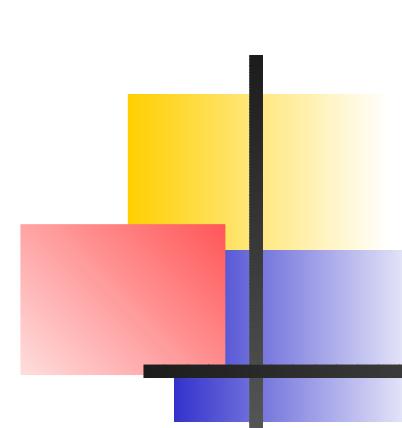
$$R = \sigma^2 I$$

$$G_i = \sigma^2 \gamma_i I_i$$

- Often several ways of writing equivalent models.

Differentiation

- $l_R = -(\log|C| + \log|R| + \log|G| + \nu \log \sigma^2 + \mathbf{y}' \mathbf{P} \mathbf{y} / \sigma^2) / 2$
- $\partial l_R / \partial \phi_i = -(-\mathbf{C}_i \mathbf{C}^{Z'Z} + 0 + \text{tr}(\mathbf{G}_i \mathbf{G}^{-1}) + 0 - \mathbf{y}' \mathbf{P} \mathbf{y}_i / \sigma^2) / 2$
 $\mathbf{y}_i = \mathbf{Z} \mathbf{G}_i \mathbf{G}^{-1} \tilde{\mathbf{u}}$
- $\partial l_R / \partial \kappa_j = -(-\mathbf{C}_j \mathbf{C}^{-1} + \text{tr}(\mathbf{R}_j \mathbf{R}^{-1}) + 0 + 0 - \mathbf{y}' \mathbf{P} \mathbf{y}_j / \sigma^2) / 2$
 $\mathbf{y}_j = \mathbf{R}_j \mathbf{R}^{-1} \tilde{\boldsymbol{\eta}}$



Two forms

- $V = \sigma^2(R + ZGZ')$
default for univariate single site analyse.
 R defined as a correlation matrix
 G defined as a variance ratio
- $V = R + ZGZ'$
used for multivariate and multisite analyses.

REML

■ Let $C = \begin{pmatrix} X'R^{-1}X & X'R^{-1}Z \\ Z'R^{-1}X & Z'R^{-1}Z + G^{-1} \end{pmatrix}$

$$P = R^{-1} - R^{-1}WC^{-1}W'R^{-1}$$

$$\begin{aligned} l_R = & -(\log|C| + \log|R| + \log|G| + \\ & \nu \log\sigma^2 + \mathbf{y}'P\mathbf{y}/\sigma^2)/2 \end{aligned}$$

Mixed model equations

■ Mixed model

$$\begin{aligned} \mathbf{y} &= \mathbf{X}\boldsymbol{\tau} + \mathbf{Z}\mathbf{u} + \boldsymbol{\epsilon} \\ \begin{pmatrix} \mathbf{X}'\mathbf{R}^{-1}\mathbf{X} & \mathbf{X}'\mathbf{R}^{-1}\mathbf{Z} \\ \mathbf{Z}'\mathbf{R}^{-1}\mathbf{X} & \mathbf{Z}'\mathbf{R}^{-1}\mathbf{Z} + \mathbf{G}^{-1} \end{pmatrix} \begin{pmatrix} \boldsymbol{\tau} \\ \mathbf{u} \end{pmatrix} &= \\ \begin{pmatrix} \mathbf{X}'\mathbf{R}^{-1}\mathbf{y} \\ \mathbf{Z}'\mathbf{R}^{-1}\mathbf{y} \end{pmatrix} \end{aligned}$$

Matrix operations

- Matrix multiplication – must be conformable

$$\begin{pmatrix} a & b & c \\ d & e & f \end{pmatrix} \begin{pmatrix} A \\ B \\ C \end{pmatrix} = \begin{pmatrix} aA + bB + cC \\ dA + eB + fC \end{pmatrix}$$

- Direct product

$$\begin{pmatrix} a \\ b \end{pmatrix} \otimes [A \ B \ C] = \begin{pmatrix} aA & aB & aC \\ bA & bB & bC \end{pmatrix}$$

Matrix operations

- Transpose

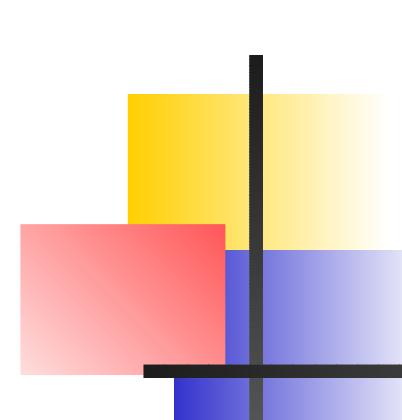
$$[a \ b \ c]' = \begin{pmatrix} a \\ b \\ c \end{pmatrix}$$

- Addition - matrices of the same order are added element by element

$$\begin{bmatrix} 1 & 2 \end{bmatrix} + \begin{bmatrix} 3 & 4 \end{bmatrix} = \begin{bmatrix} 4 & 6 \end{bmatrix}$$

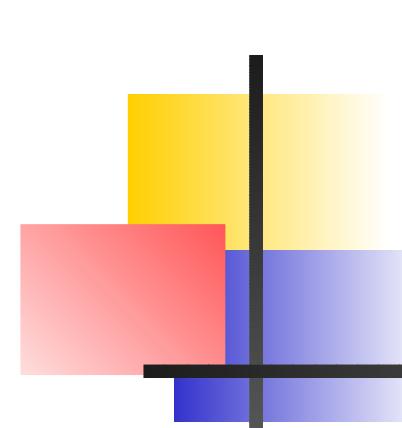
- Multiplication by scalar

$$3 \begin{bmatrix} 1 & 2 \end{bmatrix} = \begin{bmatrix} 3 & 6 \end{bmatrix}$$



Some matrices

- A *matrix* is a rectangular array of numbers
- X is the design matrix for fixed effects
 Z is the design matrix for random effects
 $W = [X \ Z]$ is the whole design matrix
 $G = \text{var}(u)$
 $R = \text{var}(\epsilon)$
 A is a relationship matrix



Some vectors

- A *vector* is a column of numbers
 - y is the response variable
 - $\hat{\tau}$ is the fixed effects
 - \tilde{u} is the random effects
 - $\tilde{\epsilon}$ is the residuals



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B1 A matrix refresher

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