

1) The constraint based version that often arises in a frequentist setting: minimize $\|X\beta - y\|_2^2$ s.t. $\|\beta\|_1 \leq t$

This constraint corresponds to the Lagrange multiplier equation

$$\underset{\beta}{\text{Argmin}} (\|X\beta - y\|_2^2 + \lambda \|\beta\|_1), \text{ where } \lambda \text{ is typically found via}$$

some cross-validation procedure or prior knowledge. Note this is the form that arises naturally in a Bayesian setting assuming a double exponential prior. Lasso can be implemented in the constraint based form using a technique like LARS, LEAST Angle Regression.

b) The covariates in this model are LDL, heart rate, ...

$$\text{let the design matrix } X = \begin{pmatrix} \text{---} X_1 \text{---} \\ \vdots \\ \text{---} X_n \text{---} \end{pmatrix} = \begin{pmatrix} LDL_1, \dots \\ \vdots \\ LDL_n, \dots \end{pmatrix}$$

Solve for β : $\underset{\beta}{\text{Argmin}} (\|X\beta - y\|_2^2 + \lambda \|\beta\|_1)$, use cross-validation for λ

c) WRT λ you could use a cross-validation scheme

d) You could link the response via $\mu = 1/(1 + \exp(-X\beta))$ using logistic regression: $Y \sim \text{Bin}(\mu)$

e) You can use L_2 penalized regression, this assumes covariates are correlated and the matrix $X^T X$ is ill-conditioned.

2) Random effects refer to population specific effects that don't represent population averages (fixed effects) ②

b) Genetic background is a random effect modelled as a random polygenic term.

c) $\langle FDR \rangle = FP / (FP + TP)$, Power = $p(T(x) \in R | H_0 = \text{False})$
 $R = \text{reject region}$
 $T(x) = \text{Test statistic}$

d) When the causatives are in the data: The power of MM and MLMM increases sharply then levels out, this sharp increase becomes slower (reverses for LM) as h^2 increases. LM follows a straight line for $h^2 = .25$ then the increase in power becomes more sensitive the increase in FDR.

When causatives dropped from data has a similar effect to increasing h^2 . LM becomes more sensitive to FDR for large FDR. MM / MLMM power responds to FDR more linearly

3) b) $p(z, x) = \prod_{i=1}^n \pi_i^{z_i} \mathcal{N}(x | \mu_i, \Sigma_i) \prod_{i=1}^n (1 - \pi_i)^{1-z_i} \exp(-\frac{1}{2}(x - \mu_i)^T \Sigma_i^{-1}(x - \mu_i))$ ($\pi_2 = 1 - \pi_1, \pi_1 = \pi$)
 $= \exp(\sum_i z_i \ln(\pi_i) - \frac{1}{2}(x - \mu_i)^T \Sigma_i^{-1}(x - \mu_i))$, thus $\prod_i p(x_i, z_i) =$
 $= L(\theta; x, z) = \exp\left[\sum_i z_i \ln(\pi_i) - \frac{1}{2}(x - \mu_i)^T \Sigma_i^{-1}(x - \mu_i)\right]$

$Q(\theta | \theta^t) = \int_z L(\theta; x, z) p(z | x, z) dz$, (I like to think of this as marginalizing out z , not computing an Expectation.)

We can compute the posteriors as: $\tau_{ij}^t = P(z_i = 1 | x_i = x_i, \theta^t) =$

$Q(\theta | \theta^t) = \sum_j \sum_i \tau_{ij}^t \left(z_i \ln(\pi_i) - \frac{1}{2}(x_i - \mu_i)^T \Sigma_i^{-1}(x_i - \mu_i) \right) \pi_i^t \mathcal{N}(x_i | \mu_i^t, \Sigma_i^t)$

n observations

* Missing $\frac{z_i}{2} \ln(\Sigma_i^{-1})$ term in all exponentials above

$\pi_i^t \mathcal{N}(x_i | \mu_i^t, \Sigma_i^t) + (1 - \pi_i^t) \mathcal{N}(x_i | \mu_i^t, \Sigma_i^t)$

3 b) $d_{\theta} Q(\theta/\theta^*) = 0 \rightarrow \begin{cases} d_{\pi} Q(\theta/\theta^*) = 0 & (1) \\ d_{\mu} Q(\theta/\theta^*) = 0 & (2) \\ d_{\Sigma} Q(\theta/\theta^*) = 0 & (3) \end{cases}$

(1) implies $\text{arg max}_{\pi} (Q(\theta/\theta^*)) = \text{arg max}_{\pi} \left(\left[\sum_i \tau_{i1}^t \right] \log \pi + \left[\sum_i (1 - \tau_{i1}^t) \right] \log (1 - \pi) \right)$

is equivalent to the MLE for the binomial

thus $\pi_i^t = \frac{\sum_i \tau_{ij}^t}{\sum_i (\tau_{ij}^t + (1 - \tau_{ij}^t))} = \frac{1}{n} \sum_i \tau_{ij}^t$

$\text{arg max}_{\mu, \Sigma} (Q(\theta/\theta^*)) = \text{arg max}_{\mu, \Sigma} \sum_i \tau_{i1}^t \left(-\frac{1}{2} \ln |\Sigma| - \frac{1}{2} (x_i - \mu_1)^T \Sigma^{-1} (x_i - \mu_1) \right)$

This is the MLE equation for multivariate gaussian (weighted)

so $\mu_1^{\text{ML}} = \frac{\sum_i \tau_{i1} x_i}{\sum_i \tau_{i1}}, \quad \Sigma_1^{\text{ML}} = \frac{\sum_i \tau_{i1} \left(\bigotimes_{j=2}^p (x_i - \mu_1)_j \right)}{\sum_i \tau_{i1}}$

μ_2, Σ_2 same as above, with 2 replacing 1

\bigotimes is the outer product between vectors, this is what I use in my R code

c) EM gives a generative model and finds the elliptical shape, K-means is faster but can't use information about the shape of the clusters.