

# Discussion on “Value of information” by Gianluca Baio (University College, London)

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The information contained in a sample is an elusive notion. The setting where the value of information is defined is the Cost–Effectiveness Analysis. The elements of CEA are:

- The net benefit of a treatment  $t$ ,

$$z(e, c|R) = R \times e - c$$

where  $e$  is the random effectiveness of the treatment,  $c$  is the random cost, and  $R$  is a deterministic quantity whose meaning is the quantity the health provider is able to pay for the unit of effectiveness.

- The distribution  $P_t(z|\theta, R)$ , is the reward of treatment  $t$ , and  $\theta$  an unknown parameter.
- The utility function of  $z$  utilized by Gianluca is

$$U(z) = z,$$

and hence the expectation  $\mathbb{E}_{z|\theta, R}U(z) \equiv NB_t(\theta|R)$ , is

$$NB_t(\theta|R) = R \mathbb{E}_{e|\theta}e - \mathbb{E}_{c|\theta}c.$$

## Expected Value of Partial Perfect Information (EVPPI).

1.- The parameter is split as  $\theta = (\phi, \psi)$  and,

$$\pi(\theta) = \pi(\phi)\pi(\psi|\phi).$$

Then, the EVPPI is given by

$$\text{EVPPI} = \mathbb{E}_{\phi} \max_t \mathbb{E}_{\psi|\phi} NB_t(\theta|R) - \max_t \mathbb{E}_{\theta} NB_t(\theta|R) \geq 0.$$

- Is there a subparameter  $\phi$  of interest in CEA?
- It is not the expectation of the net benefit the quantity of interest?

2.- Which the the role of  $R$  in EVPPI?

- 3.- For a given  $R$  and the set of competing treatments  $\mathcal{T} = \{T_t, t = 1, \dots, k\}$ , the treatment given by

$$\max_{t=1, \dots, k} \mathbb{E}_{\theta} NB_t(\theta, R)$$

is an optimal treatment with respect to the rewards

$$P_t(z|R) = \int P_t(z|\theta, R)\pi(\theta) d\theta, t = 1, \dots, k.$$

- The treatment defined by

$$\mathbb{E}_{\phi} \max_t \mathbb{E}_{\psi|\phi} NB_t(\theta|R),$$

- is a treatment in  $\mathcal{T}$  ?
  - Which are the rewards?
- 4.- The parameters are always integrated out w.r.t. a prior. No data on  $e$  and  $c$  are used.
- 5.- I would like to have some properties of the EVPPI.

## Expected value of sampling information (EVSI)

1.- EVSI is defined for  $x = (e, c)$  with distribution  $P(x|\theta)$  as

$$\text{EVSI} = \left| \mathbb{E}_x \max_t \int NB_t(\theta|R)\pi(\theta|x) d\theta - \max_t \int NB_t(\theta|R)\pi(\theta)d\theta \right|$$

The treatment in  $\{T_t, t = 1, \dots, k\}$  from

$$\max_t \int NB_t(\theta|R)P_t(x|\theta)\pi(\theta) d\theta$$

depends on  $x$ . Then, is the treatment defined by

$$\int \left[ \max_t \int NB_t(\theta|R)P_t(x|\theta)\pi(\theta) d\theta \right] dx$$

in the class?

Remember that you assume that if the optimal treatment does not change the value of the information is zero.

## Expected value of sampling information (EVSI)

- 2.- Again, the definition of EVSI does not need data.
- 3.- The above expressions are computed from simulations, for instance, from the model

$$NB_t(\theta) = \mathbb{E}_{\psi|\phi} NB_t(\theta) + \varepsilon, \quad \varepsilon \sim N(\varepsilon|0, \sigma^2).$$

I have difficulties to understand this model.

**Thanks Gianluca for your work**

# Discussion on “Bayes–optimal frequentist intervals and tests”

by Peter Hoff (Duke University)

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- Peter considers heterogenous independent normal sampling models

$$N(y_j|\theta_j, \sigma^2) \quad j = 1, \dots, p,$$

where  $\theta_1, \dots, \theta_p$  are i.i.d. with prior distribution

$$N(\theta_j|\mu, \tau^2), \quad j = 1, \dots, p.$$

- $\mathbf{y}_j = (y_{j1}, \dots, y_{jn_j})$  is a sample of  $y_j$  of size  $n_j$ .
- The interest is on estimating  $\theta_j$ .
- $\sigma, \mu, \tau$  are nuisance parameters for which no prior is assigned.



- The likelihood of  $\sigma$ ,  $\mu$ ,  $\tau$  for the data  $\mathbf{y} = (\mathbf{y}_1, \dots, \mathbf{y}_p)$ , is

$$\ell(\sigma, \mu, \tau; \mathbf{y}) = \prod_{j=1}^p \int \left[ \prod_{i=1}^{n_j} N(y_{ji} | \theta_j, \sigma^2) \right] N(\theta_j | \mu, \tau^2) d\theta_j$$

and  $\hat{\sigma}^2$ ,  $\hat{\mu}$  and  $\hat{\tau}^2$  maximizes the likelihood. All data are here.

- An “empirical” Bayesian estimator of  $\theta_j$  is the posterior expectation plus MLE,

$$\hat{\theta}_j = \frac{n_j / \hat{\sigma}^2}{n_j / \hat{\sigma}^2 + 1 / \hat{\tau}^2} \bar{y}_j + \frac{1 / \hat{\tau}^2}{n_j / \hat{\sigma}^2 + 1 / \hat{\tau}^2} \hat{\mu}.$$

- The MLEs do not bother me as they do a very good job for moderate or large sample sizes.

## Groups uncertainty

- The distribution  $N(\theta_j|\mu, \tau^2)$  represents the uncertainty of  $\theta_j$  around  $\mu$  in group  $j$ , but it does not inform us about the uncertainty across groups.
- However, the smaller the  $\hat{\tau}^2$  the bigger the influence of  $\hat{\mu}$  on the estimator  $\hat{\theta}_j$ .
- Further, it is obvious that heterogeneity disappears if  $\tau = 0$ . Thus, for testing homogeneity we would testing the null hypothesis

$$H_0 : \tau = 0,$$

against

$$H_1 : \tau \geq 0.$$

- After testing we can proceed with either the homogeneous model (if  $H_0$  is accepted) or the heterogeneous one (if  $H_1$  is accepted)

- A difficulty arises when using  $p$ -values for testing:  
We cannot compute the probability of  $H_0$  and  $H_1$ , and hence we do not know the weights to be given to the inference (estimation, etc.) from the null model

$$\left\{ N(y|\theta, \sigma^2), \pi(\theta) \right\}$$

and from the alternative model

$$\left\{ N(y_j|\theta_j, \sigma^2), N(\theta_j|\mu, \tau^2) \quad j = 1, \dots, p \right\}.$$

## Heterogeneity

Another related question is whether we should restrict ourself to fully heterogeneity (we note that partial heterogeneity is deemed possible)

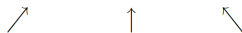
Can we cluster some of the sampling models?

- Use clustering based on Bayesian model selection of the partition models (Hartigan 1990, Casella et al. 2014).
- The uncertainty on the cluster models must be incorporated into the final inference.

# Bayesian meta-analysis scheme

Meta-model

$$f(x|\theta), \pi^N(\theta)$$



Linking

distribution

$$\pi(\theta_1|\theta)$$

...

$$\pi(\theta_k|\theta)$$



Experimental  
Models

$$f(x_1|\theta_1), \pi^N(\theta_1)$$

...

...

$$f(x_j|\theta_j), \pi^N(\theta_j)$$

...

...

$$f(x_p|\theta_p), \pi^N(\theta_p)$$

►  $x$  is a latent variable (non observable)

## The prior distributions

- As the experimental model and the meta-model are the same,  $\theta_j$  and  $\theta$  are identically distributed with distribution  $\pi^N(\cdot)$
- The linking distribution,  $\pi(\theta_i|\theta)$  has to be compatible with  $\pi^N(\cdot)$ , that is

$$\int \pi(\theta_j|\theta)\pi^N(\theta) d\theta_j = \pi^N(\theta), \quad \int \pi(\theta_j|\theta)\pi(\theta) d\theta = \pi^N(\theta_j)$$

- $\pi(\theta_i|\theta)$  should be able to accommodate different degrees of concentration of  $\theta_i$  around  $\theta$

## A general linking distribution

From the model comparison between a generic experimental model

$$f(x|\theta_j), \pi^N(\theta_j)$$

and the meta-model for a fixed  $\theta$

$$f(x|\theta)$$

the intrinsic prior  $\pi^I(\theta_j|\theta, t)$  (Berger and Pericchi 1996, Moreno et al. 1998) satisfies:

- i)  $\pi^I(\theta_j|\theta, t)\pi^N(\theta)$  has marginals  $\pi^N(\theta)$  and  $\pi^N(\theta_j)$ .
- ii) Through  $t$  we accommodate a given degree of concentration of  $\theta_i$  around  $\theta$

**Example.** Incidence data of wound infection in cesarean section for patients treated with prophylactic antibiotics versus placebo from seven randomized controlled trials (Sutton et al. 2002)

Trial	Antibiotics		Placebo	
	$n_i$	$x_i$	$n_i$	$x_i$
1	100	3	33	0
2	11	1	17	5
2	42	0	40	0
4	167	4	140	5
5	36	1	25	1
6	115	12	117	15
7	16	0	16	1

Model:  $Bin(x_i|\theta_i, n_i)$ ,  $i = 1, \dots, 7$ . We are interested in estimating  $\theta$ , the meta-parameter.



## The top clusters

Antibiotics		Placebo	
Cluster Models	Post. Prob.	Cluster Models	Post. Prob.
{1, 3, 4, 5, 7}, {2}, {6}	0.29	{1}, {2}, {3}, {4}, {5}, {6}, {7}	0.26
{1}, {2}, {3}, {4}, {5}, {6}, {7}	0.10	{1, 3}, {2}, {4}, {5}, {6}, {7}	0.04
{1, 3, 4, 5}, {2}, {6}, {7}	0.04	{1, 3, 4, 5, 7}, {2, 6}	0.03
{1, 3, 4, 5, 6, 7}, {2}	0.04	the rest	< 0.03
{1, 3, 4, 7}, {2}, {5}, {6}	0.04		
the rest	< 0.03		

Posterior expectation of the meta-incidence  
of wound infection in cesarean

Without clustering			After clustering		
Antibiotics	Placebo	Incidence reduction	Antibiotics	Placebo	Incidence reduction
0.024	0.045	-0.021	0.057	0.070	-0.013

Thanks Peter for you work