



Centre for Academic
Primary Care

NIHR | School for Primary
Care Research

How do GPs make sense of diagnostic tests?

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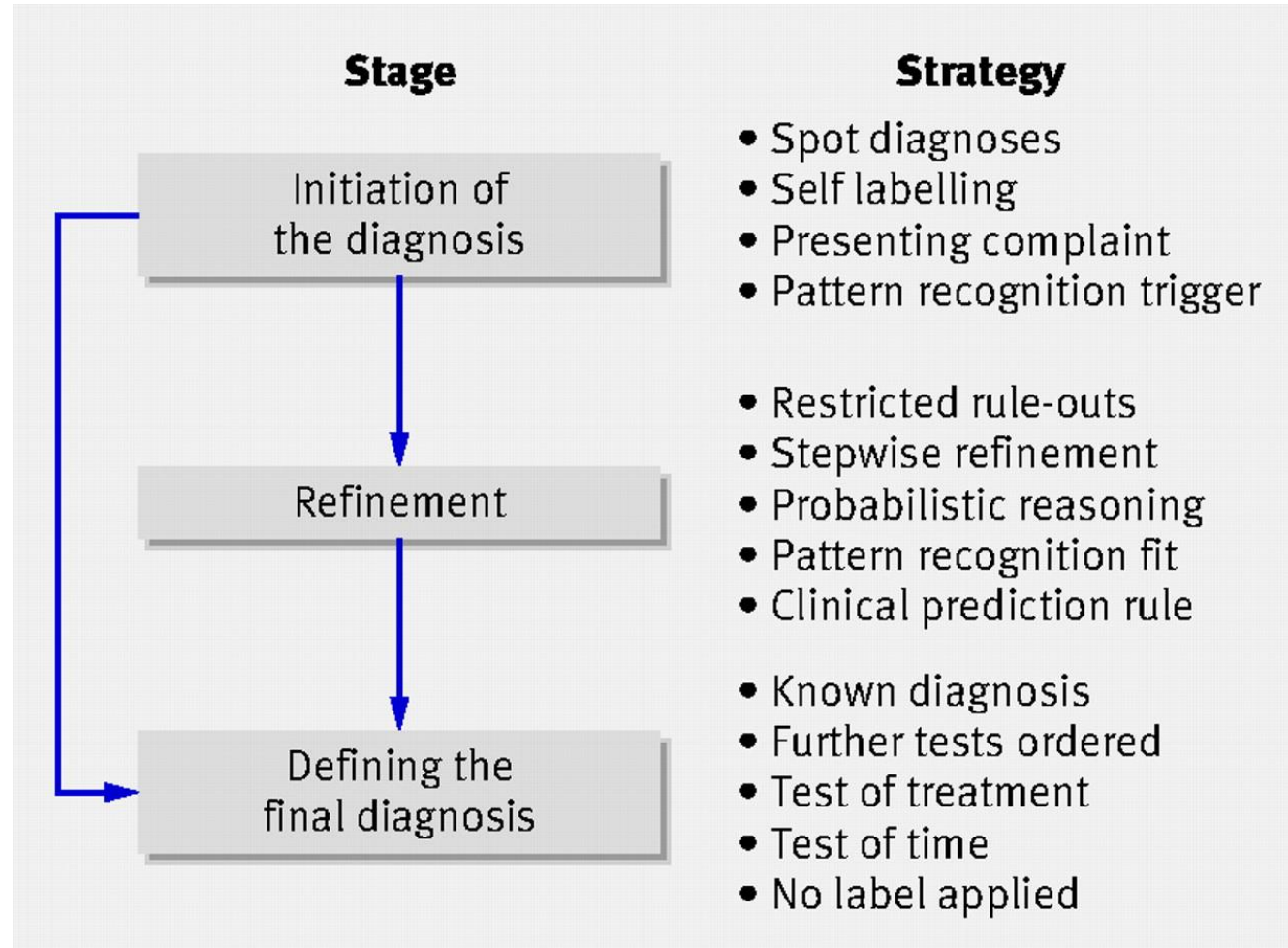
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Outline

- How do GPs make a diagnosis?
- What are diagnostic tests?
- Interpreting tests – the theory
- Interpreting tests – the reality

How do GPs make a diagnosis?



What is a diagnostic test?

“any procedure, or test, that tries to confirm or identify the presence or absence of a target condition”

- Includes laboratory tests, point of care tests, imaging, invasive procedures, patient history, physical examination, questionnaires, test of time, test of treatment

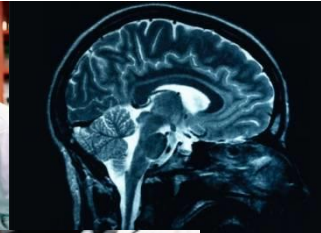
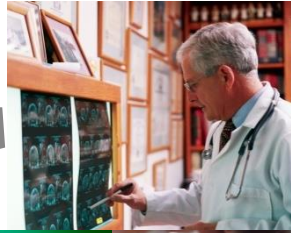
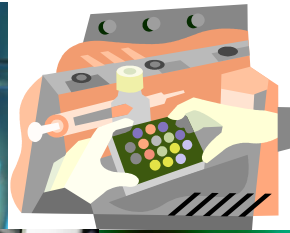


Table 11. The CAGE Questionnaire.

- Have you ever felt the need to Cut down on your drinking?
- Have you ever felt Annoyed by criticism of your drinking?
- Have you ever felt Guilty about your drinking?
- Have you ever felt the need to drink a morning Eye-opener?



What do tests do?

Tests determine if you have a disease

What do tests do?

~~Tests determine if you have a disease~~

Tests determine your chances of having a disease

What do tests do?

~~Tests determine if you have a disease~~

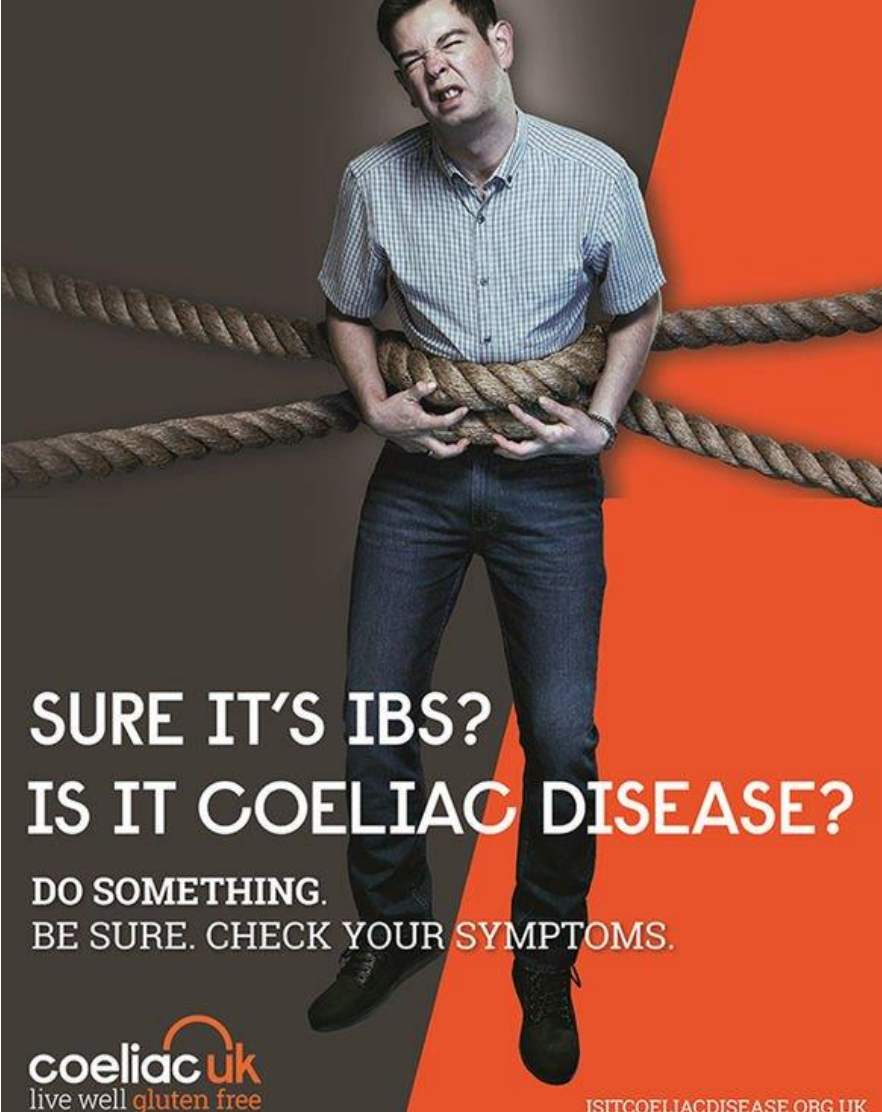
~~Tests determine your chances of having a disease~~

Tests update your chances of having a disease

Mary's story



Could Mary have coeliac disease?



**SURE IT'S IBS?
IS IT COELIAC DISEASE?**

DO SOMETHING.
BE SURE. CHECK YOUR SYMPTOMS.


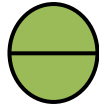

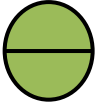
coeliacuk
live well gluten free

ISITCOELIACDISEASE.ORG.UK

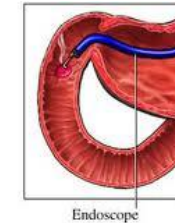
How do we calculate coeliac disease test accuracy?

Index test result
Blood test (tissue
transglutaminase)



		True disease state	
			
Index test result		True positive	False positive
		False negative	True negative

True disease state
Duodenal biopsy
(reference standard)



$$\text{Sensitivity} = \text{TP} / (\text{TP} + \text{FN})$$

$$\text{Specificity} = \text{TN} / (\text{TN} + \text{FP})$$

Interpreting diagnostic tests – the theory

Sensitivity: Proportion of individuals *with* the condition who test *positive*

$$\text{Sensitivity} = TP / (TP + FN)$$

Specificity: Proportion of individuals *without* the condition who test *negative*

$$\text{Specificity} = TN / (TN + FP)$$

Mary's story - continued

- Mary's blood test result is positive
- Sensitivity of IgA TTG is estimated at 90.7%, specificity 87.4%^{9*}
- What is the likelihood that she has coeliac disease?
- Does she need a biopsy to confirm?



⁹*Elwenspoek, et al. *Identifying the optimum strategy for identifying adults and children with coeliac disease: systematic review and economic modelling*. NIHR Journals Library.

Conditional probabilities

- **Sensitivity:** Proportion of individuals with the condition who test positive

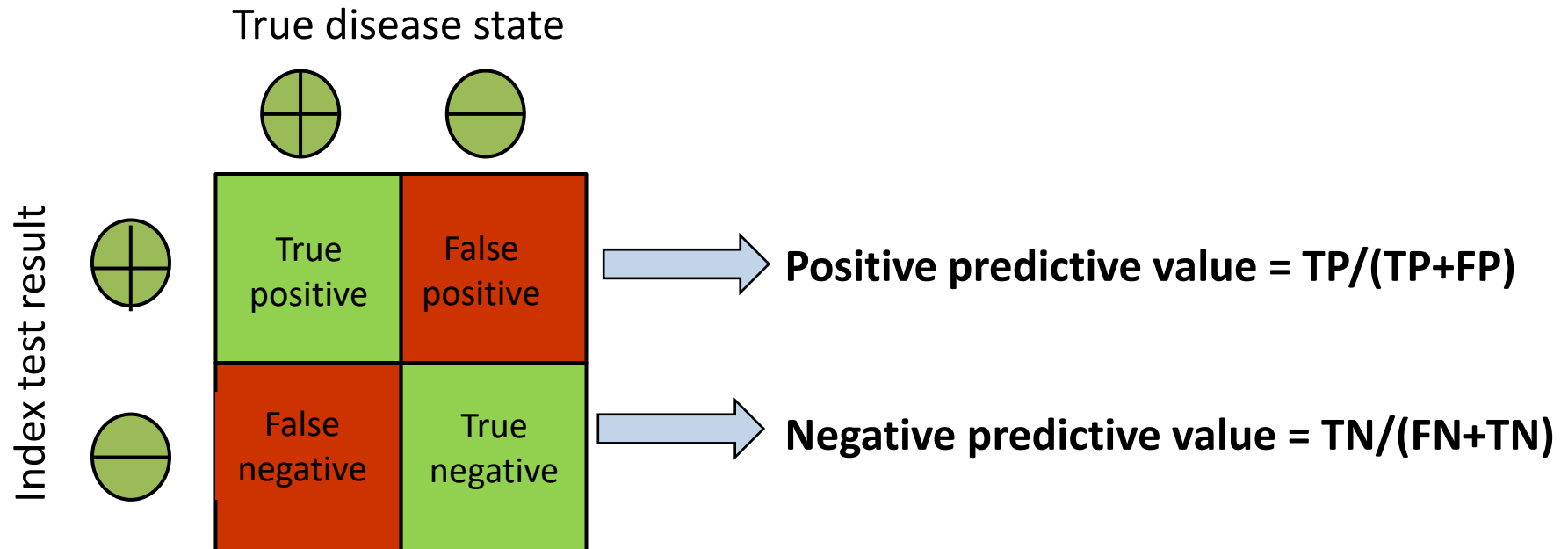
$$P(\text{Test positive}|\text{disease})$$

- **Positive Predictive Value (PPV):** Probability of having the disease in a patient with a positive result

$$P(\text{Disease}|\text{Test positive})$$

$$P(\text{Disease}|\text{Test positive}) = \frac{P(\text{Disease}) \times P(\text{Test positive}|\text{Disease})}{P(\text{Test positive})}$$

Predictive values



PPV is mathematically dependent on **pre-test probability (for an individual) or prevalence (at a population level)**

Estimating pre-test probability

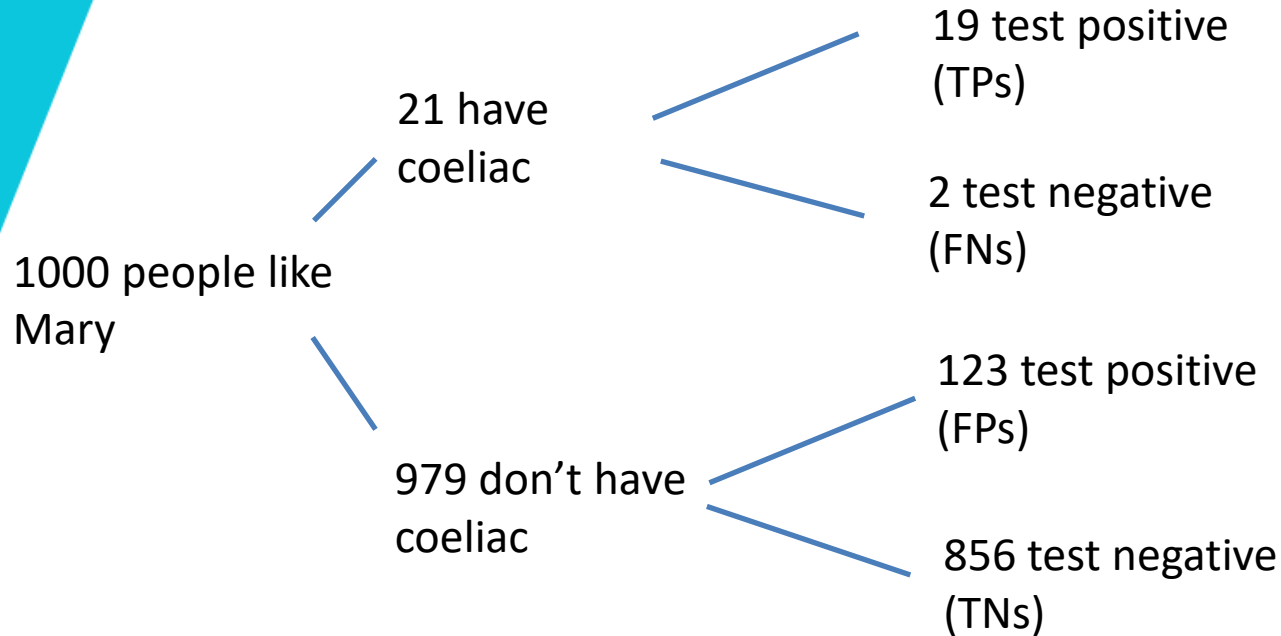
- Prevalence coeliac disease: roughly 1%
- History and examination used to update pre-test probability:
 - Symptoms? Family history? Past medical history?

The screenshot shows a web browser window with the URL <https://calculator.testingwisely.com>. The page title is "Testing Wisely Disease Risk Calculator" and it includes a "Playground Calculator" link. The interface is divided into two main sections. On the left, a dark grey box displays the "CHANCE OF Celiac disease BEFORE TEST" as "2.1 %", accompanied by a horizontal bar chart showing the percentage on a scale from 0 to 100. On the right, a white form contains two numbered steps. Step 1, "What disease are you Testing for?", has a dropdown menu set to "Celiac Disease". Step 2, "What is the pre-test probability?", is divided into three sections: "INCIDENCE" with a checked box for "General US population"; "EXPOSURE RISK" with unchecked boxes for "Type 1 DM", "1st degree relative", and "Type 1 DM and 1st degree relative"; and "SYMPTOMS" with a checked box for "Malabsorption" and unchecked boxes for "Short stature/failure to thrive", "Unexplained anemia", "Neuropathy", "Arthralgias", and "Unexplained transaminitis". At the bottom of the form, a text input field contains "2.1" with a label "*Estimated pre-test probability".

Calculating predictive values:

Sensitivity = 90.7%, specificity = 87.4%

Mary has a pre-test probability of coeliac of 2.1%



If Mary tests positive, the probability she has coeliac is:

$$\frac{19}{19 + 123} = \mathbf{13.4\%}$$

If Mary tests negative, the probability she has coeliac is:

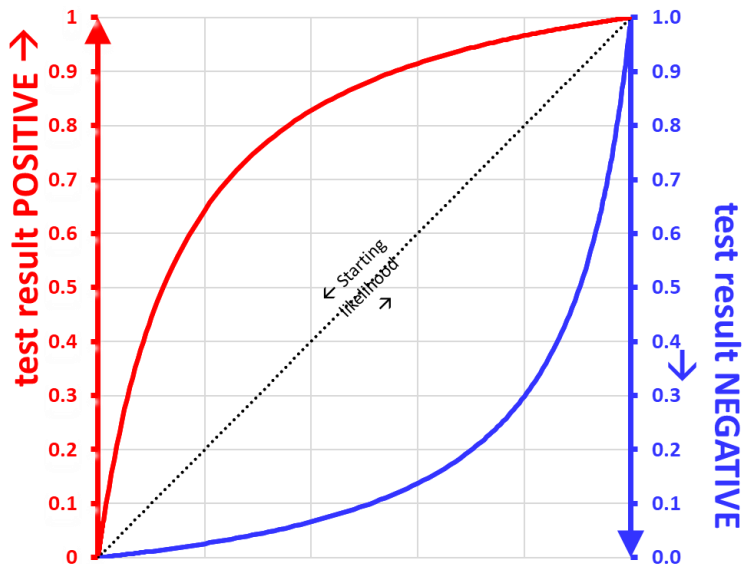
$$\frac{2}{2 + 856} = \mathbf{0.02\%}$$

Calculating predictive values: tools

$$P(A|B) = \frac{P(A) \times P(B|A)}{P(B)}$$

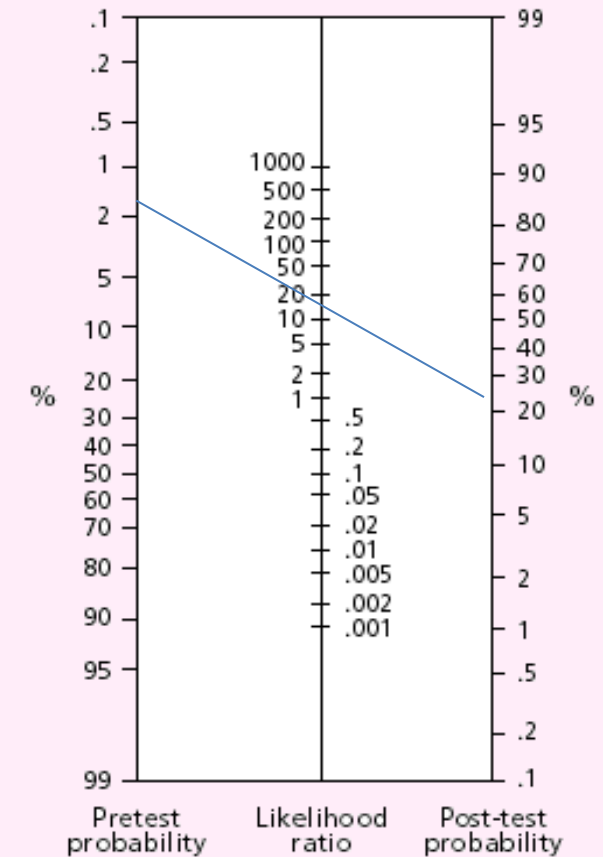
where

- $P(A|B)$ = how likely is A given B
- $P(B|A)$ = how likely is B given A
- $P(A)$ = how likely is A overall
- $P(B)$ = how likely is B overall

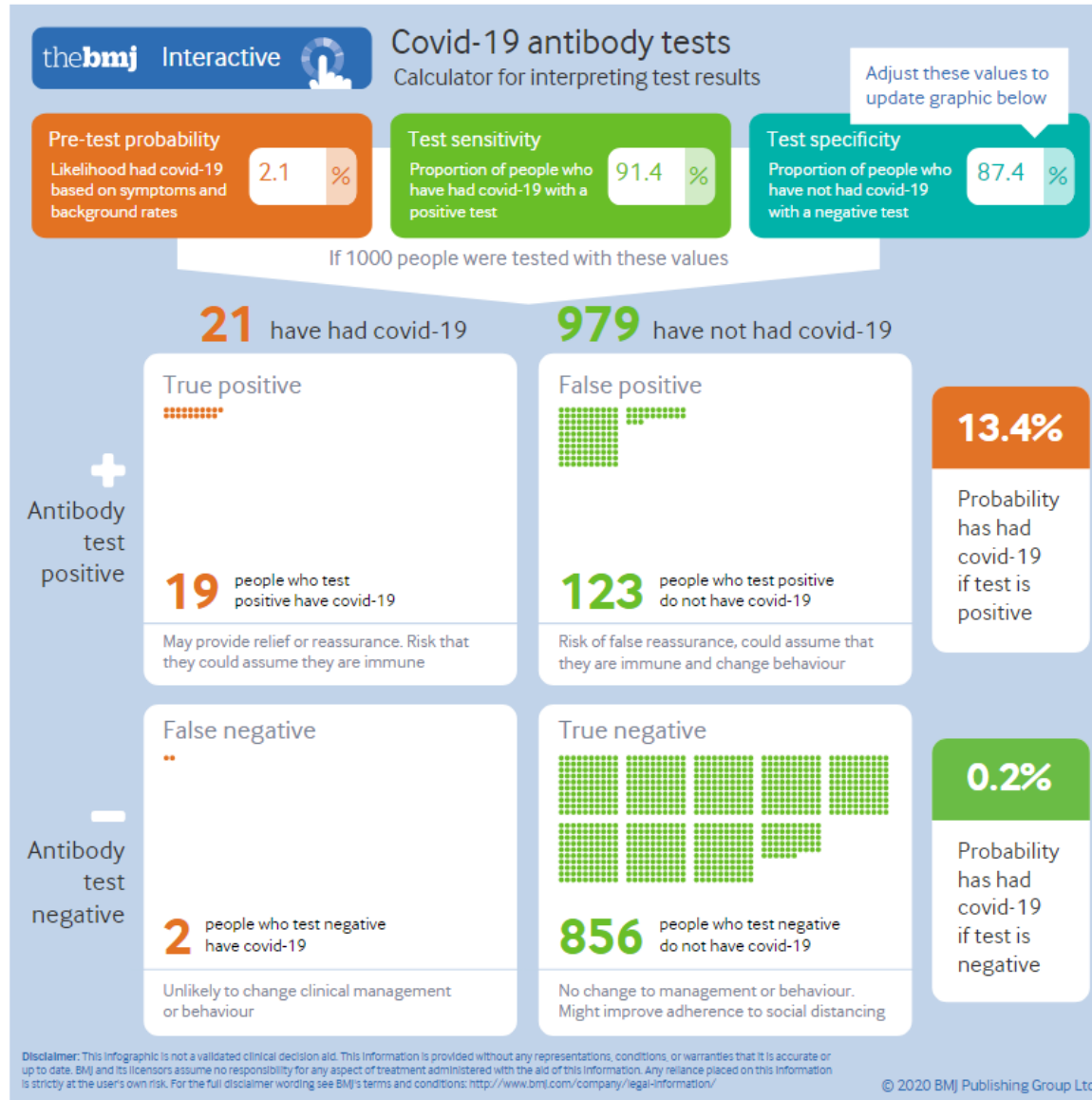


Bayes' nomogram

Pre-test probability is located on the first axis and joined to the appropriate likelihood ratio on the second axis. The post-test probability is then read off the third axis.



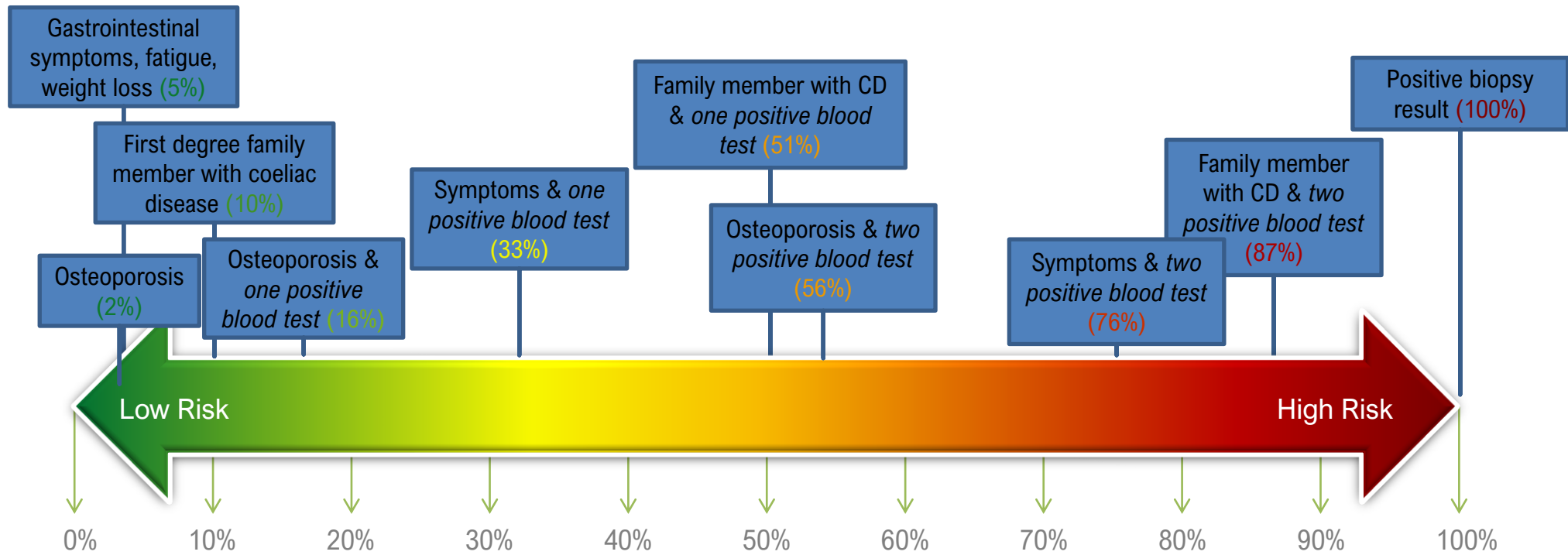
Calculating predictive values: tools



Estimating the predictive values: the reality

- Heuristics (learned mental short cut):
 - Anchoring (the pre-test probability)
 - Adjusting (based on the test result)





Does Mary need a biopsy?

- *How sure would you like to be before starting a gluten free diet?*

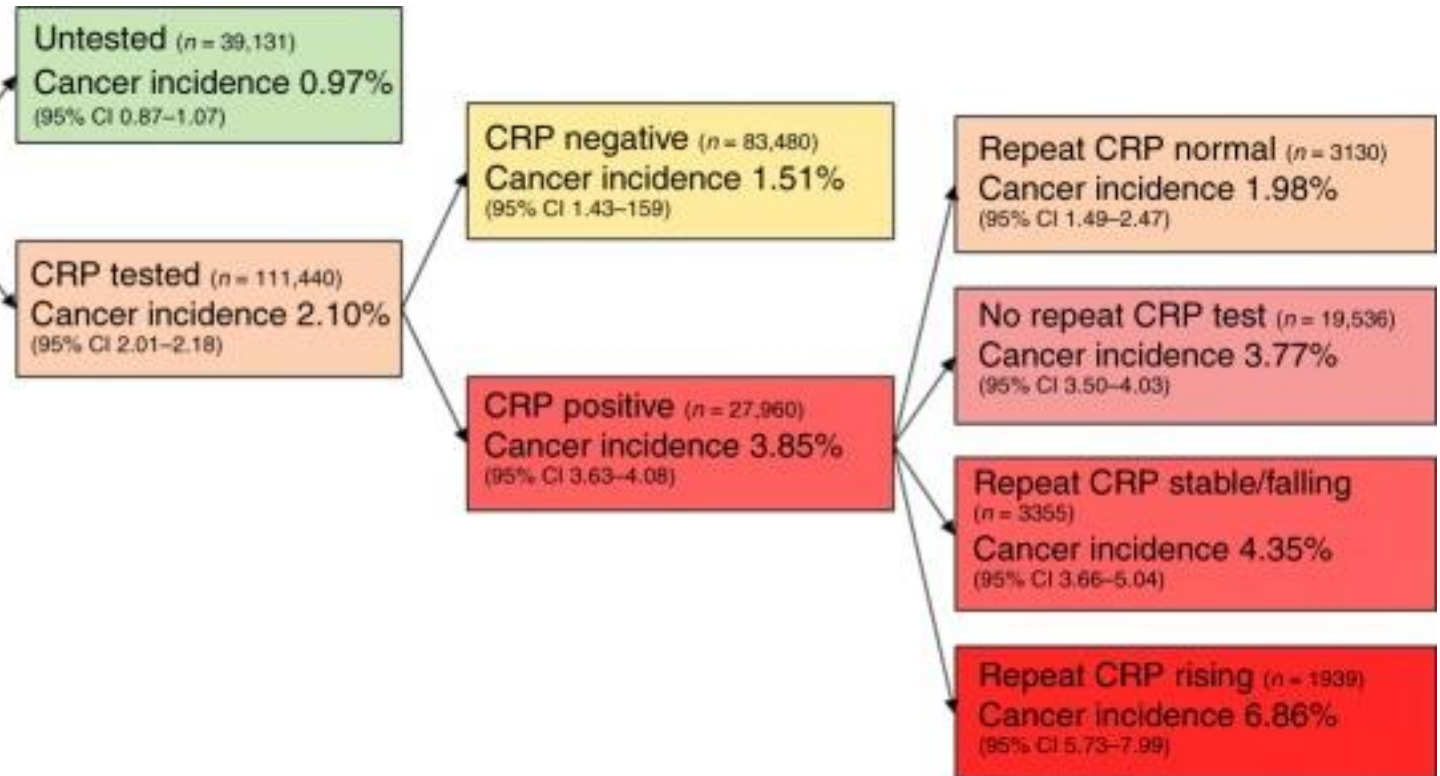


Martin's story

- Martin has the same symptoms as Mary
- His coeliac serology tests were negative
- All other blood tests were normal apart from a raised CRP test result (non-specific marker of inflammation)

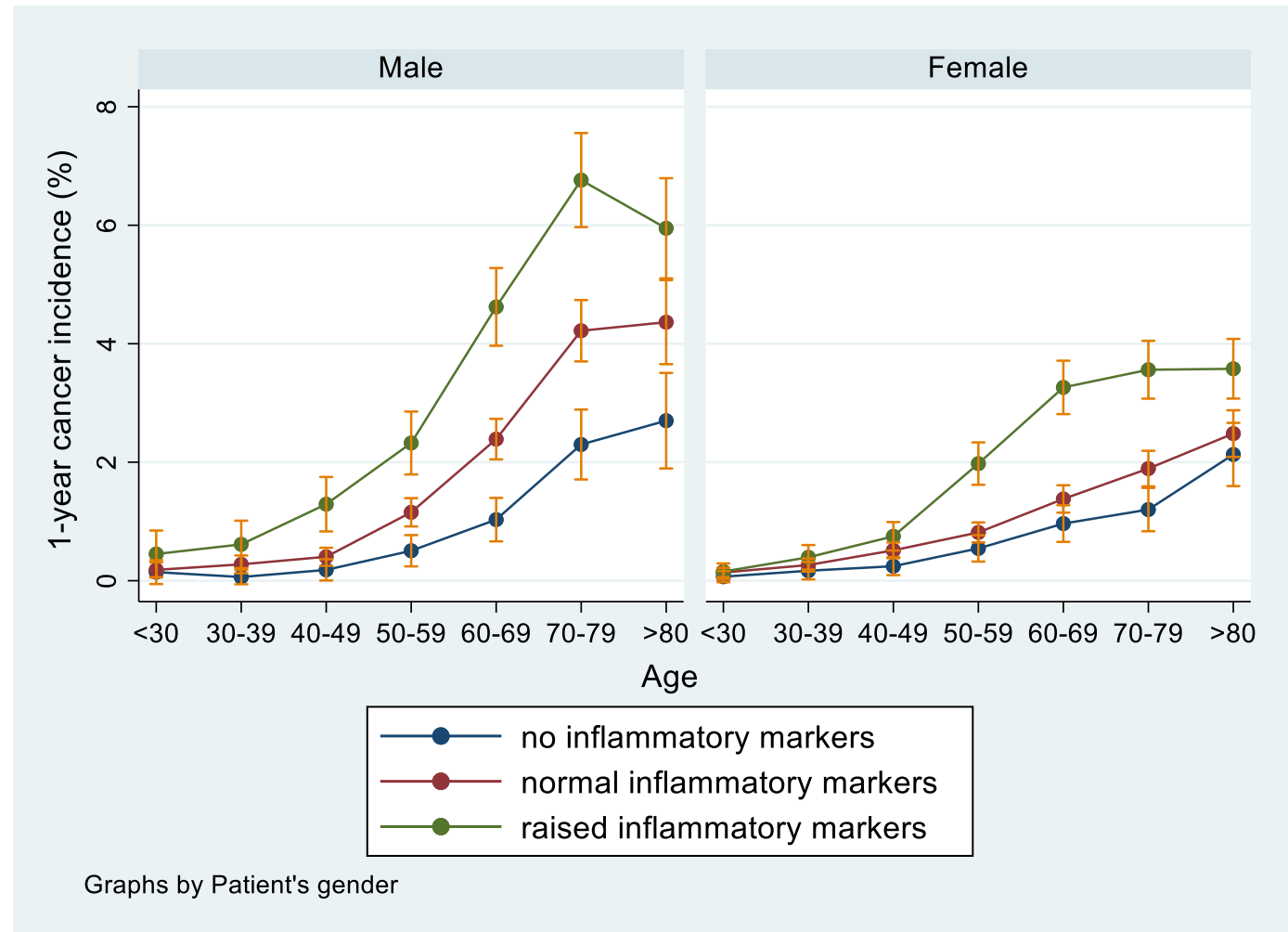


Testing is a series of Bayesian steps



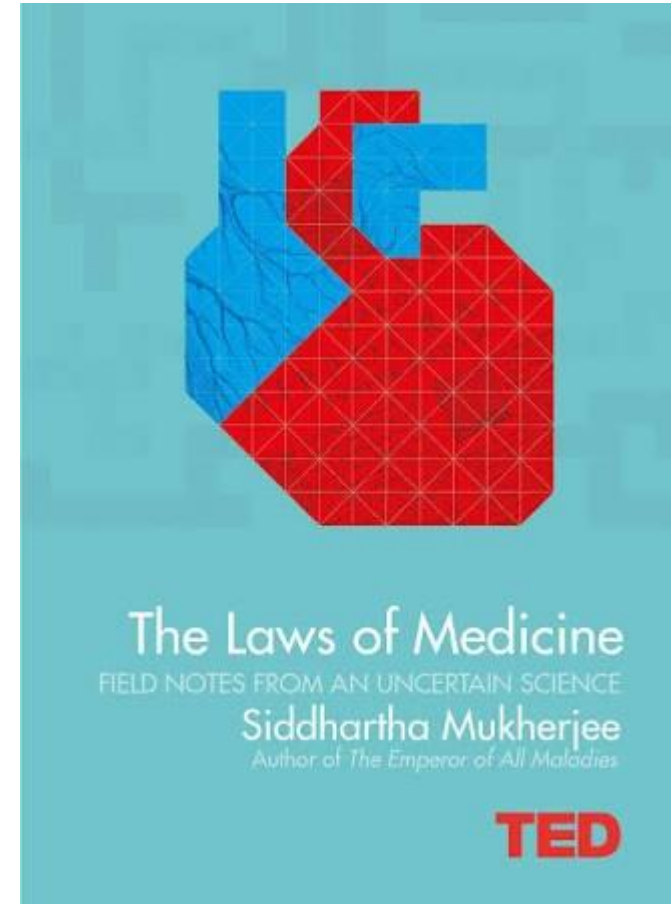
The paradox of the normal test result

- The mere fact that a test result has been performed increases the risk of cancer
- This additional risk is only partly eliminated by a negative test result



Cognitive biases – base rate neglect

- Anchoring and adjustment heuristics can be prone to bias
- Base rate neglect - the tendency to trust results of an 'objective' test more than ones own 'subjective' clinical judgement.
- “A strong intuition is much more powerful than a weak test”
 - Siddhartha Mukherjee’s first ‘Law of Medicine’



Take home messages

- Tests *update your chance* of having disease
- To understand a test result you need to know the test accuracy and the pre-test probability
- Doctors use mental shortcuts (heuristics) called anchoring and adjusting
- This can be prone to cognitive biases
- More tools to help interpret test results could help diagnostic decision making



Thank you

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