



Pogojna verjetnost in Bayesov izrek

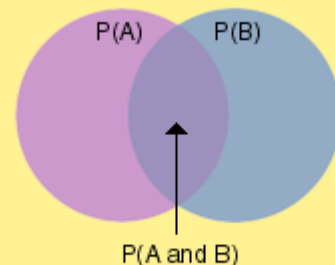
Iztok Grabnar

Univerza v Ljubljani, Fakulteta za farmacijo

Maj, 2012

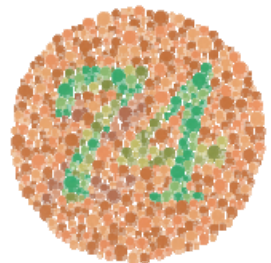
Pogojna verjetnost

- Verjetnost dogodka "a" pri pogoju, da se je zgodil dogodek "b"
- Zapis:
 - $p(a|b)$
 - | pomeni "če je", "če imamo", "če velja"
- Uporaba:
 - Diagnostični testi (občutljivost/specifičnost)
 - Analiza podatkov, primerjava modelov
 - Markovski procesi

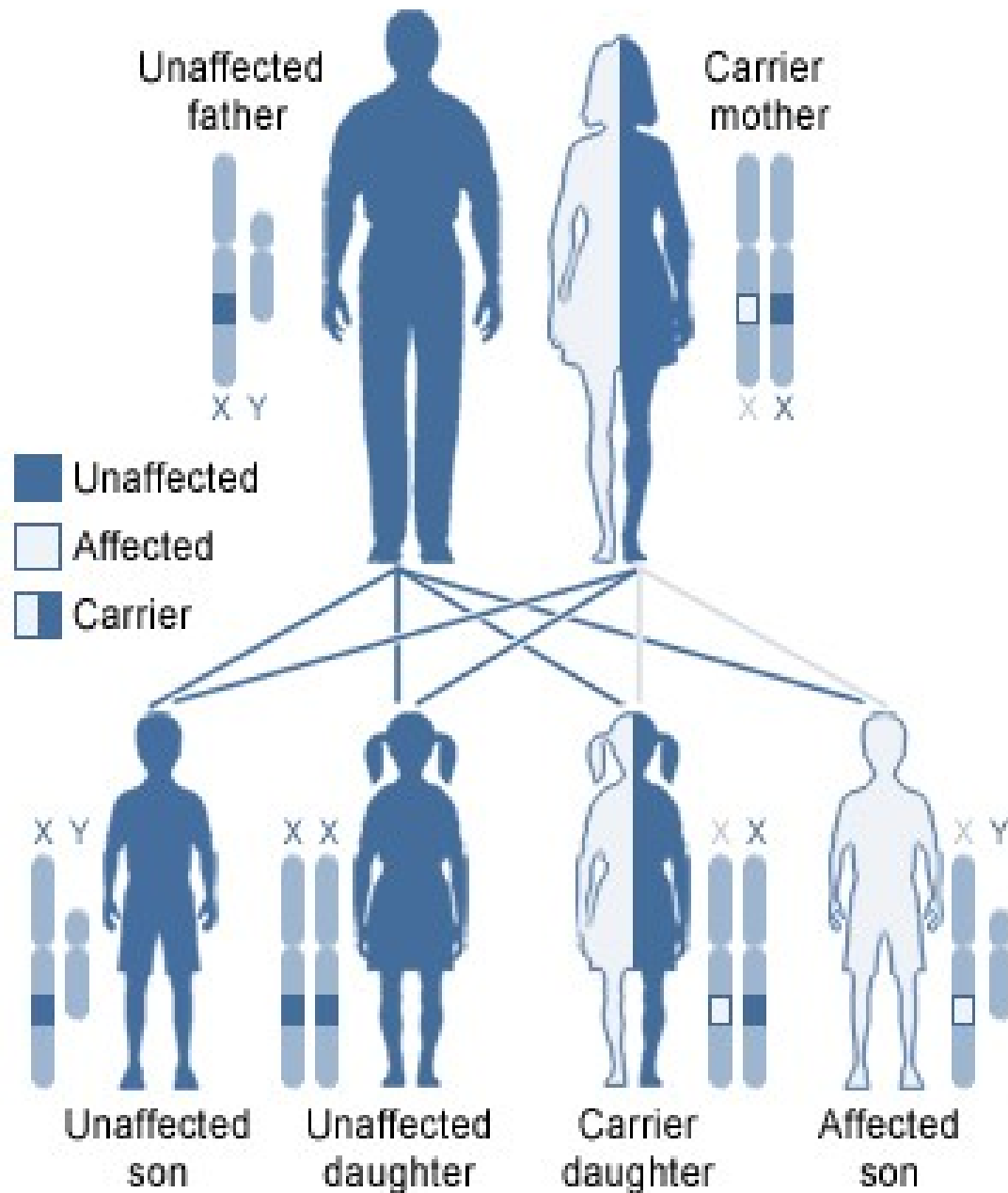


Pogojna verjetnost

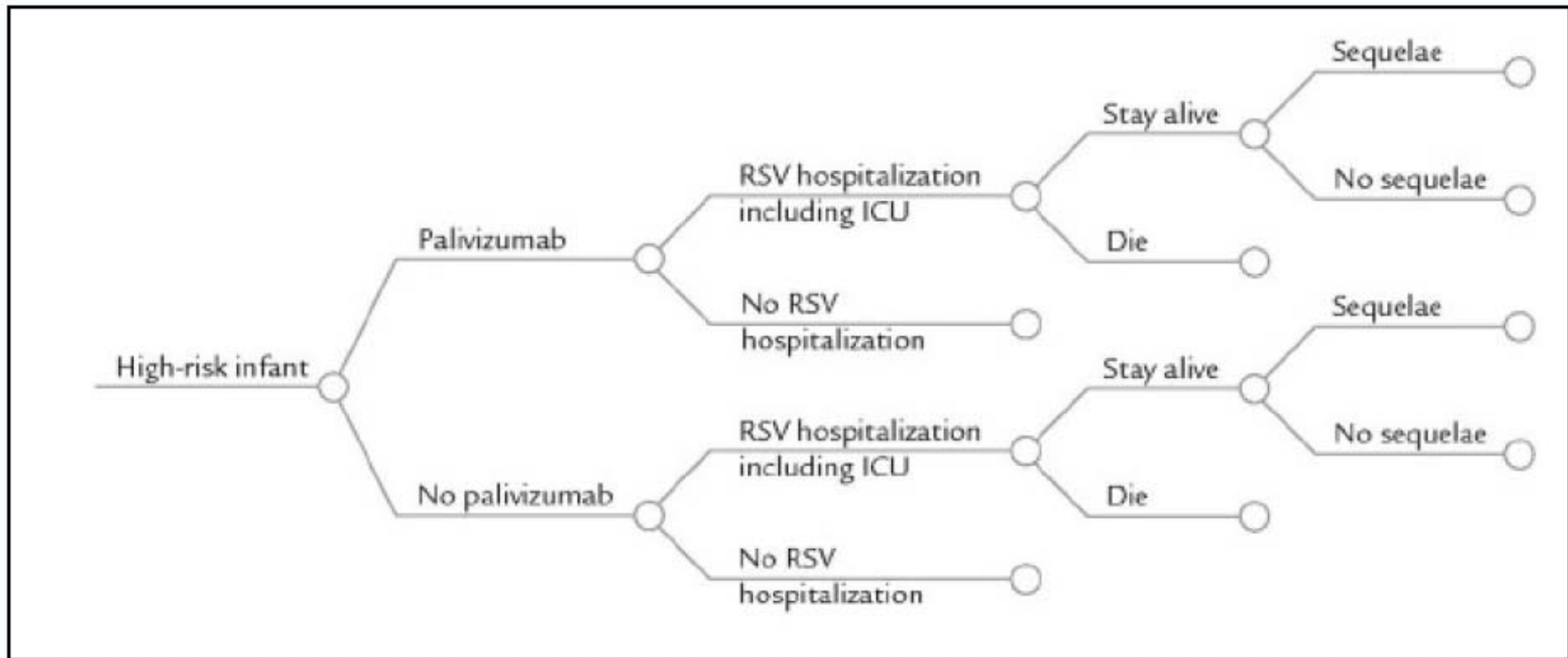
- ❑ Verjetnost, da bo otrok deček ($p=0,51$)
- ❑ Verjetnost krvne skupine A ($p=0,43$)
- ❑ Kakšna je verjetnost, da bo otrok deček s krvno skupino A?
- ❑ Verjetnost barvne slepote ($p=0,04$)
- ❑ Kakšna je verjetnost, da bo otrok deklica z barvno slepoto?



X-linked recessive, carrier mother







Odločitveni model



Slika 1: Model odločitvenega drevesa za ocenjevanje stroškovne učinkovitosti palivizumaba (ang.)

Igralne karte

			
13	13	13	13
13	13	7	13

$P(S)$

$P(F)$

$P(S|F)$

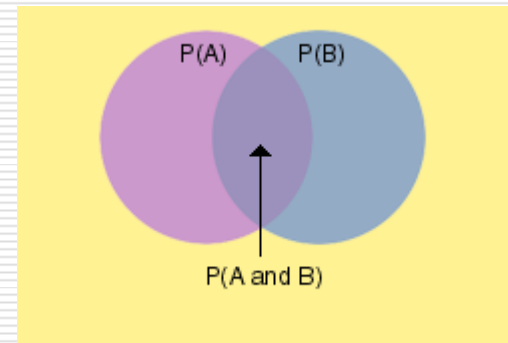
$P(F|S)$

Primer pogojne verjetnosti

- Diagnostični klinični test
- Dogodka
 - Dogodek A: Pacient ima bolezen
 - Dogodek B: Test je pozitiven

□ Zapis:

- $p(A \cap B)$ ■ Verjetnost, da ima pacient bolezen in pozitiven izid testa
- $p(A \cap B')$ ■ Verjetnost, da ima pacient bolezen, a je izid testa negativen (**lažno negativen**)
- $p(A' \cap B)$ ■ Verjetnost, da pacient nima bolezn, a je izid testa pozitiven (**lažno pozitiven**)
- $p(A|B)$ ■ Verjetnost, da ima pacient bolezen, če je izid testa pozitiven
- $p(A|B')$ ■ Verjetnost, da ima pacient bolezen, če je izid testa negativen



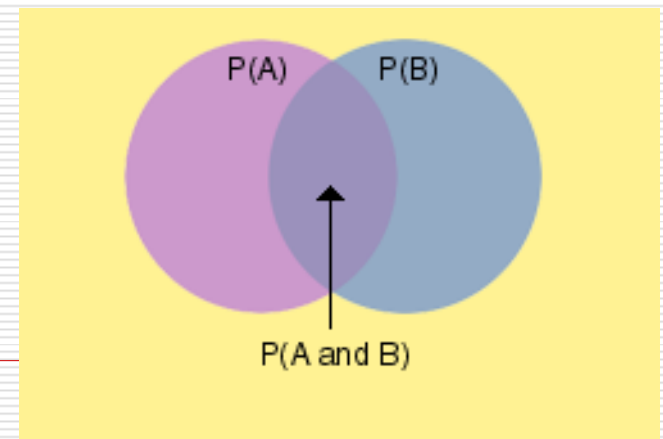
30% pacientov, ki so bili v letu 2008 sprejeti v enoti nujne medicinske pomoči UKC Ljubljana zaradi bolečine v trebušni votlini, je imelo akutno vnetje slepiča. 70% bolnikov z vnetjem slepiča je imelo telesno temperaturo nad 37,5°C. Pri pacientih, ki niso imeli vnetja slepiča, pa je bila telesna temperatura višja od 37,5°C pri 40% bolnikov.

-
- Kaj lahko rečemo o dogodku A, če poznamo izid B?
 - Pozitiven test = bolezen, negativen test = zdrav?
 - Diagnostični testi niso idealni!

$$p(A \cap B) = p(A|B)p(B) = p(B|A)p(A)$$
$$p(A|B') = \frac{p(A \cap B')}{p(B')} = \frac{p(A \cap B')}{1 - p(B)}$$

$$p(A|B) = \frac{p(B|A)p(A)}{p(B)} = \frac{p(B|A)p(A)}{p(B|A)p(A) + p(B|A')p(A')}$$

- Odvisnost dogodkov A in B
 - Ali je $p(A)$ odvisna od $p(B)$?
 - Ali je $p(B)$ odvisna od $p(A)$?



Bayesov teorem



Thomas Bayes (1702-1761)

- Dogodki A_i kjer je $i = 1$ do k so:
 - Nezdružljivi: $A_i \cap A_j = 0$ za vse i, j
 $P(A_1 \cup \dots \cup A_k) = 1$

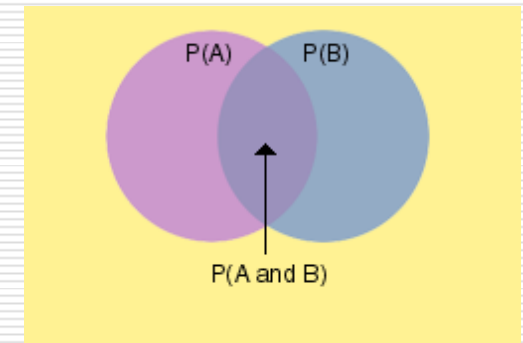
- Za vsak dogodek B

$$p(B) = p(B|A_1)p(A_1) + \dots + p(B|A_k)p(A_k)$$

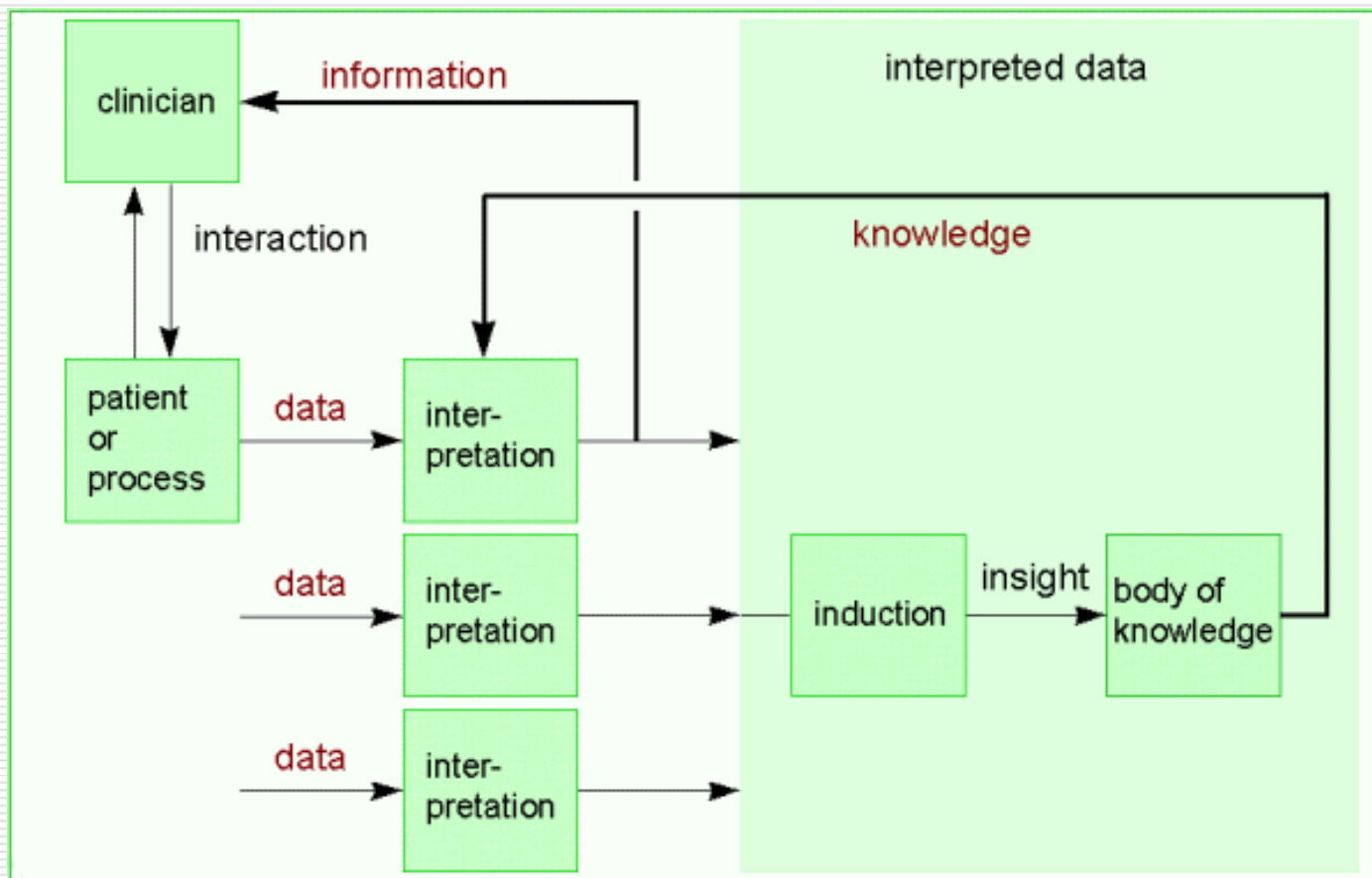
$$p(B) = \sum_{i=1}^k p(B|A_i)p(A_i)$$

- Bayesov teorem

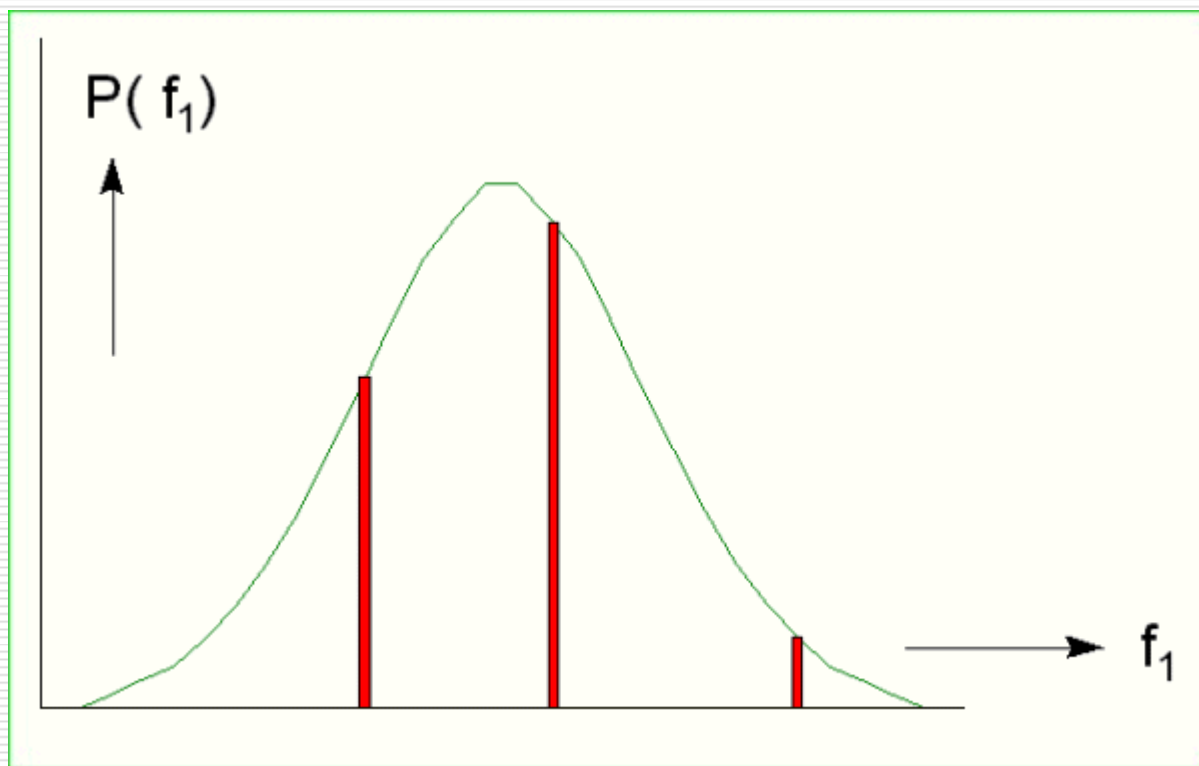
$$p(A_j|B) = \frac{p(A_j \cap B)}{p(B)} = \frac{p(B|A_j) \cdot p(A_j)}{\sum_{i=1}^k p(B|A_i)p(A_i)}$$

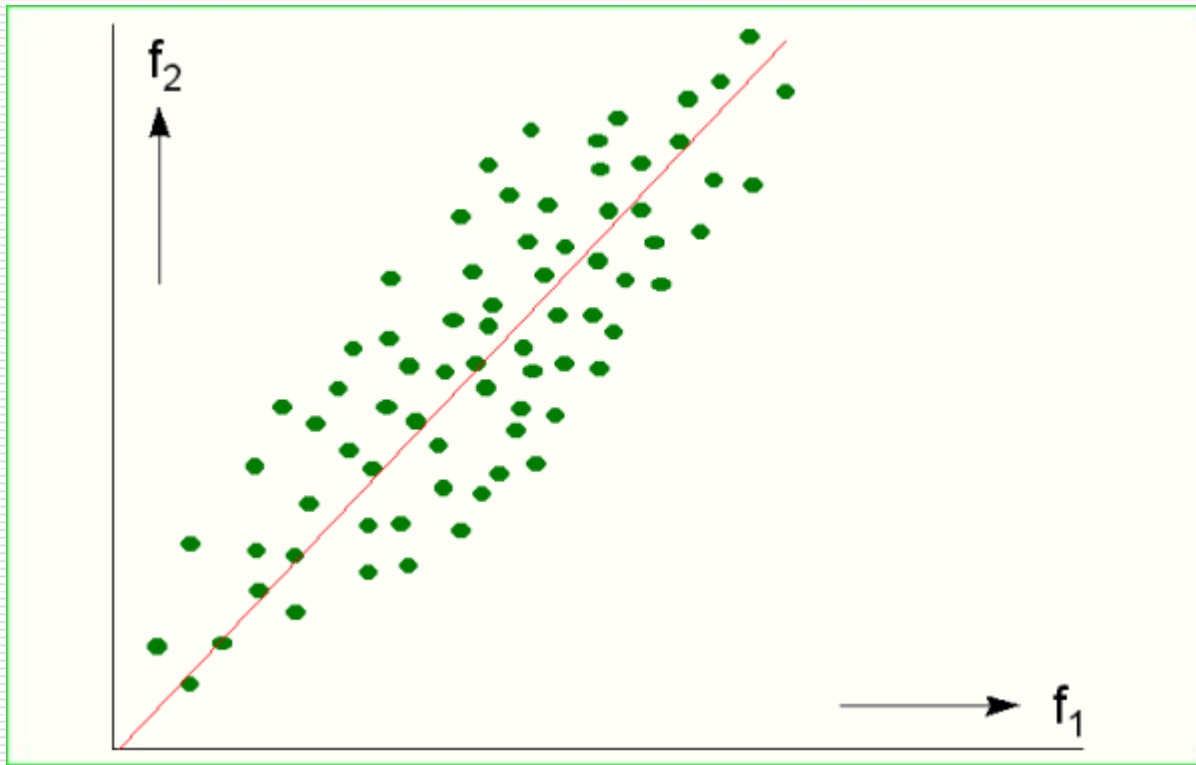


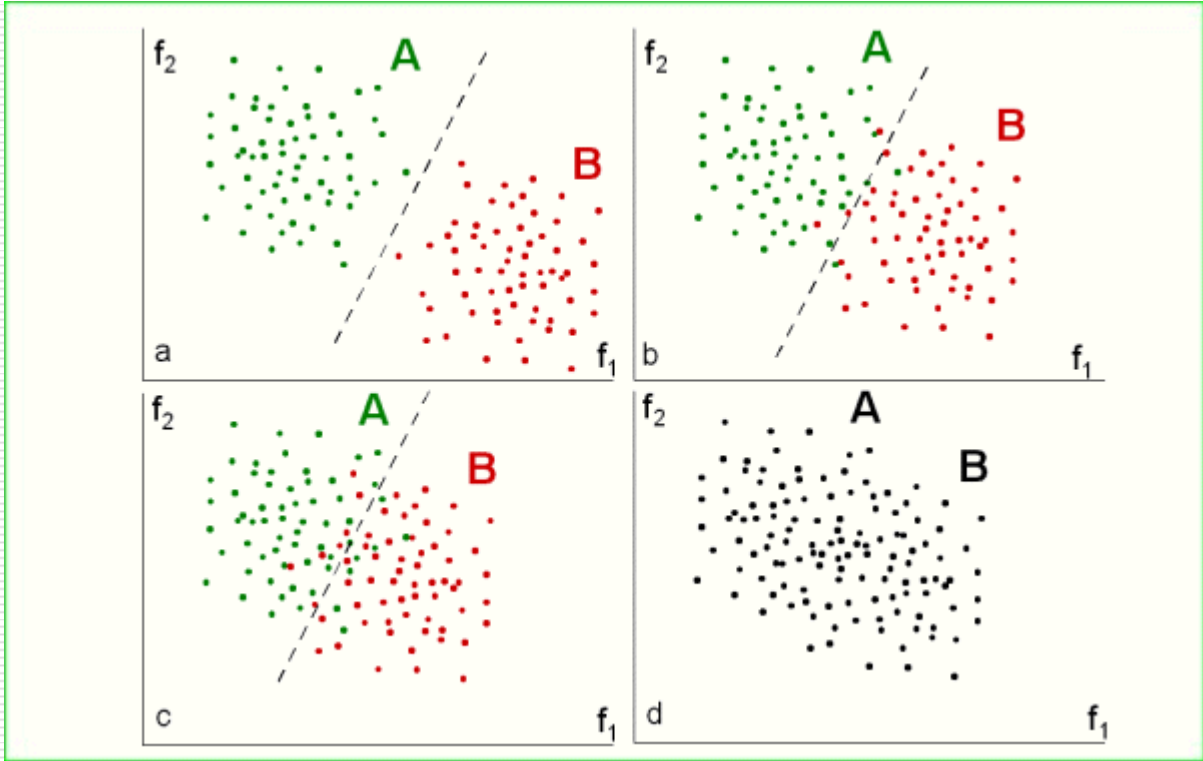
Podatek, informacija in znanje

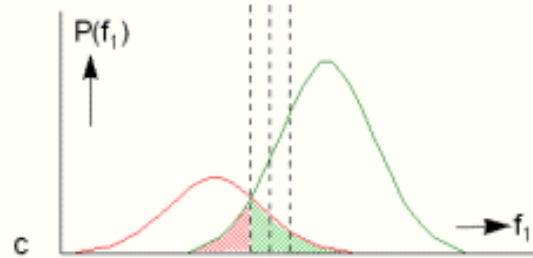
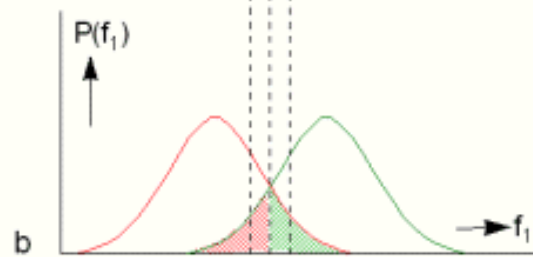
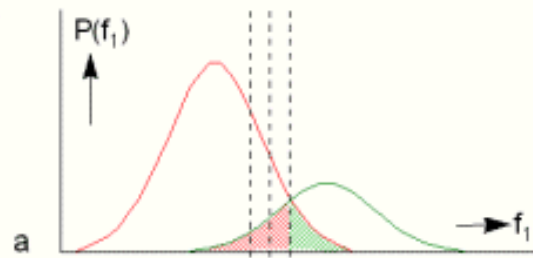


Učenje









15.01 Relationships between True Positive (TP), True Negative (TN), False Positive (FP), and False Negative (FN)

		Decision Model		
		+	-	
Truth	+	TP	FN	100%
	-	FP	TN	100%

Table 15.1. Relationships between True Positive (TP), True Negative (TN), False Positive (FP), and False Negative (FN).

Občutljivost in specifičnost

15.02 Illustration of Sensitivity, Specificity, and Predictive Value

		Decision Model		
		+	-	
Truth	+	a	b	a + b
	-	c	d	c + d
		a + c	b + d	a + b + c + d

Table 15.2. Illustration of Sensitivity, Specificity, and Predictive Value (see text).

Občutljivost $TP = a / (a + b)$

Specifičnost $TN = d / (c + d)$

Pozitivna napovedna vrednost $PNV = a / (a + c)$

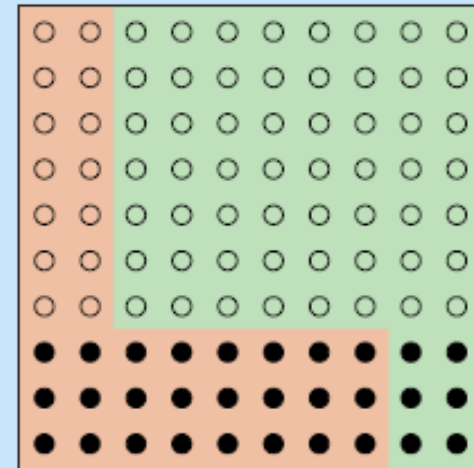
Negativna napovedna vrednost $NNV = d / (b + d)$

Understanding sensitivity and specificity with the right side of the brain

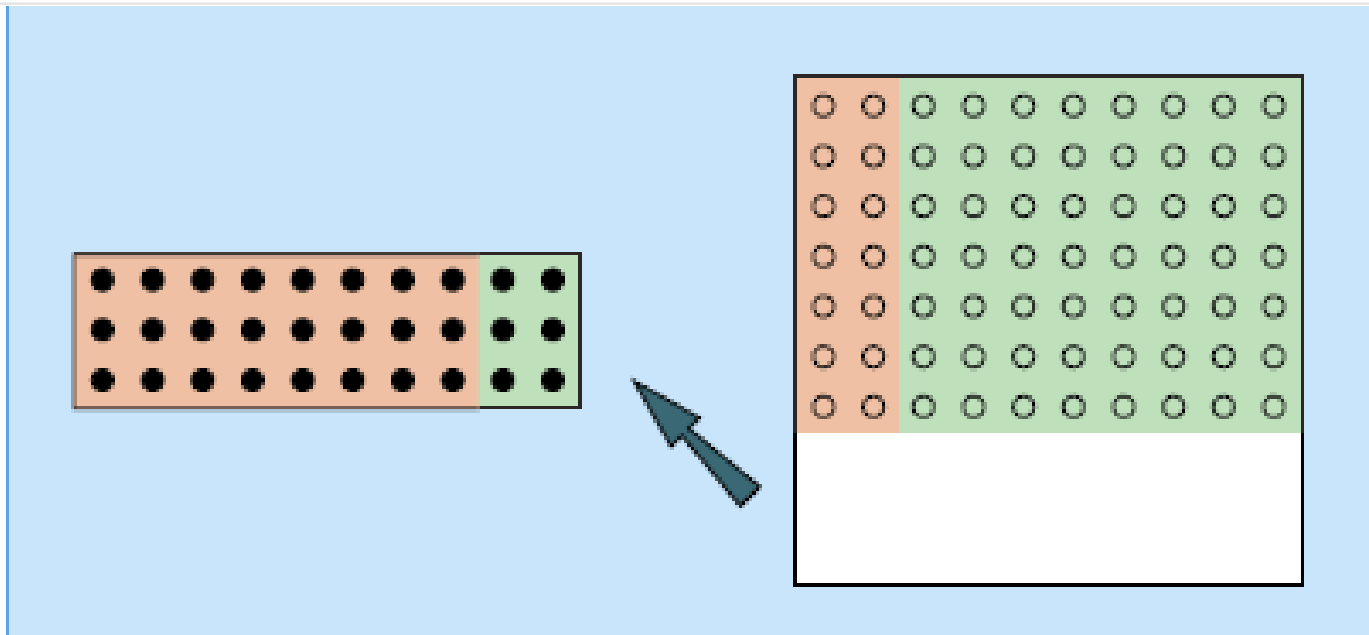
-is a well person
-is a person with a disease
-is a negative test result
-is a positive test result

and therefore....

- □is a well person who tests negative (a true negative)
- □is a person with a disease who tests positive (a true positive)
- □is a well person who tests positive (a false positive)
- □is a person with a disease who tests negative (a false negative)

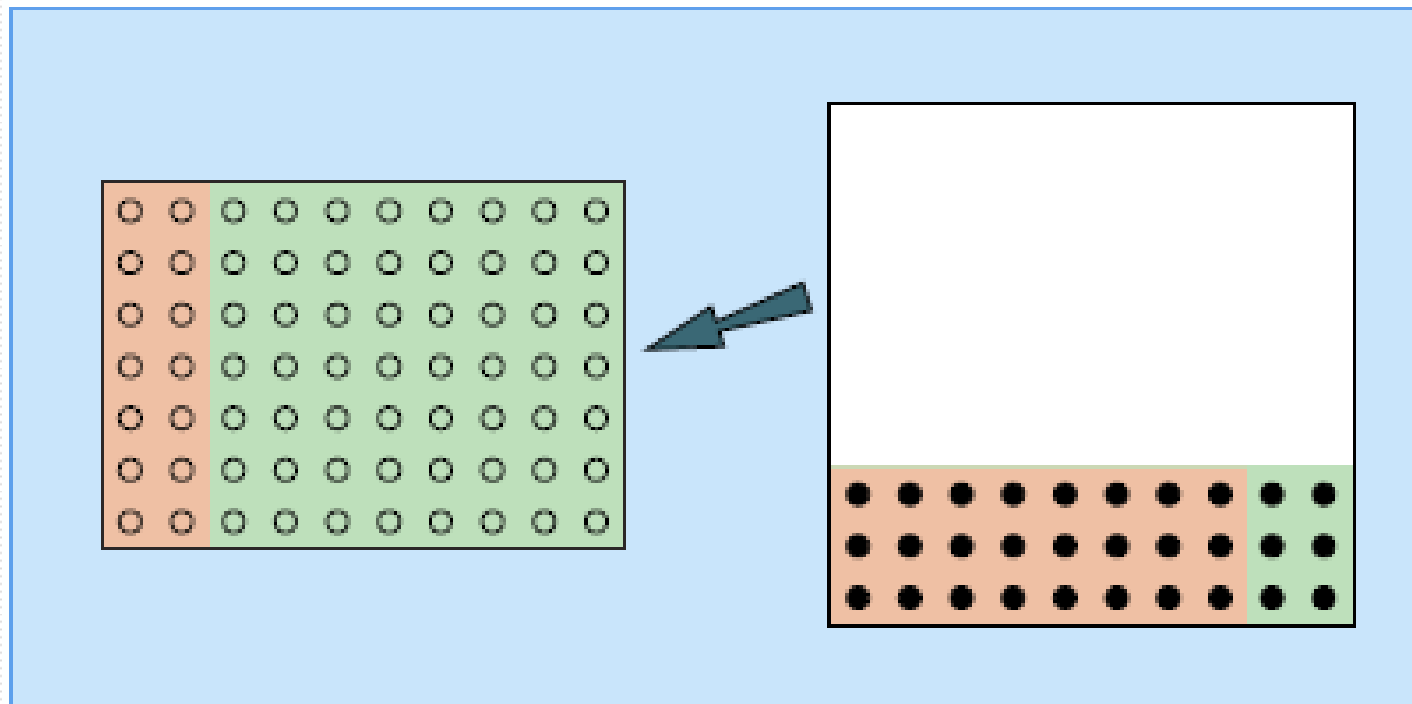


Občutljivost



$$24/30=80\%$$

Specifičnost



$$56/70 = 80\%$$

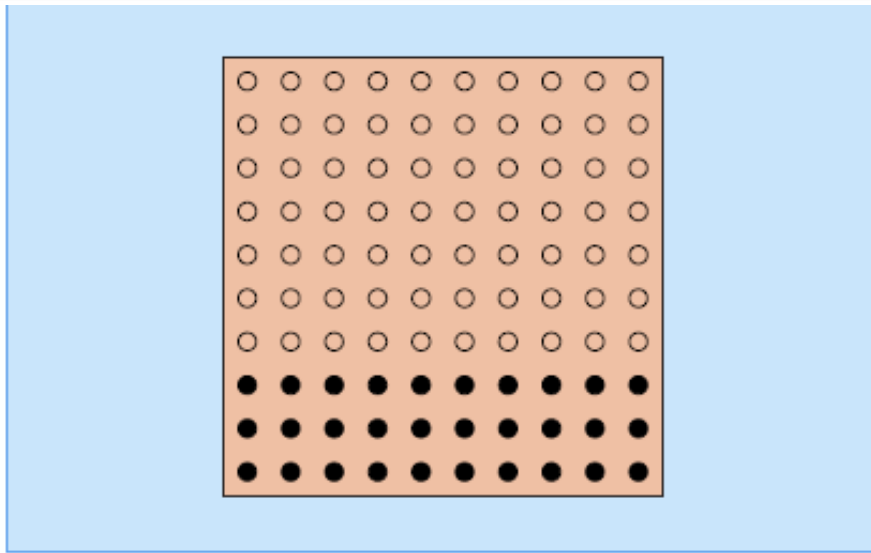


Fig 6 Test with 100% sensitivity

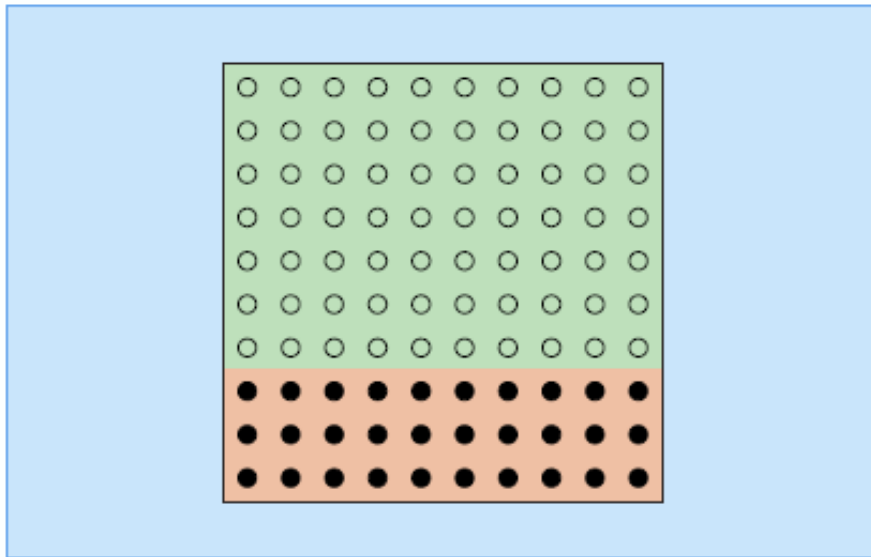
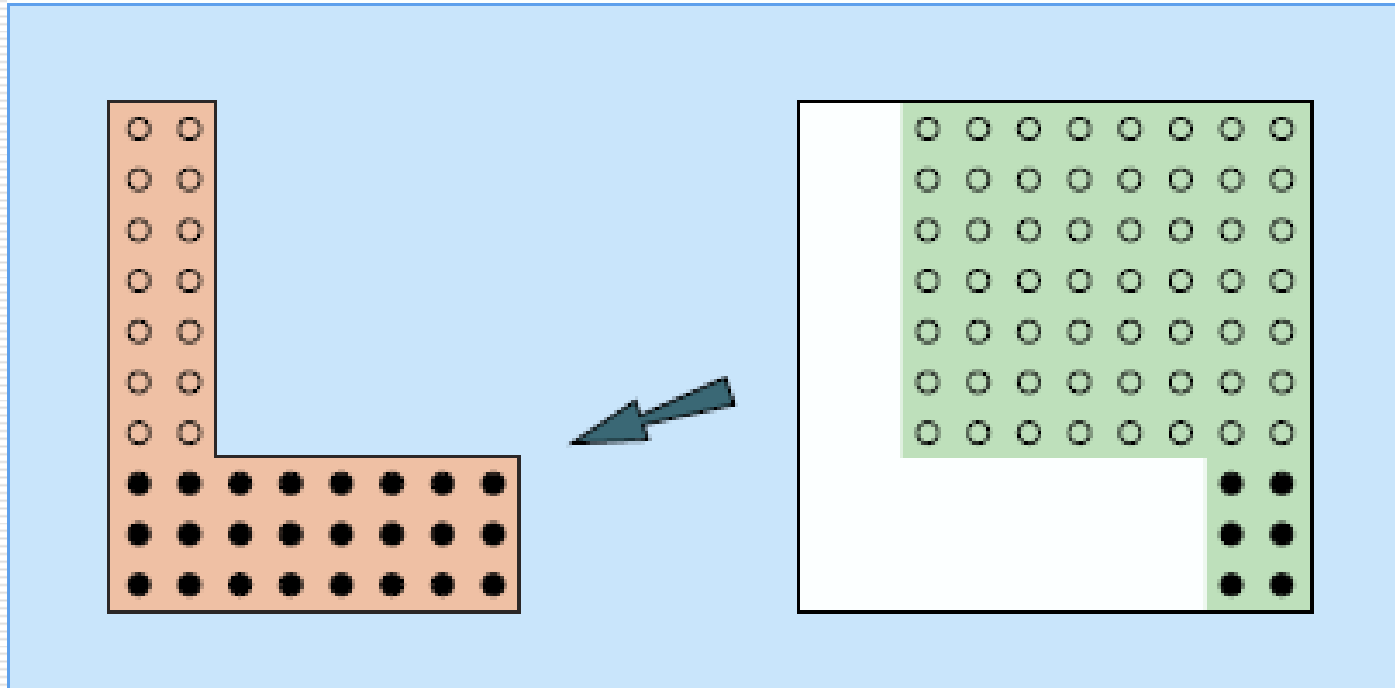


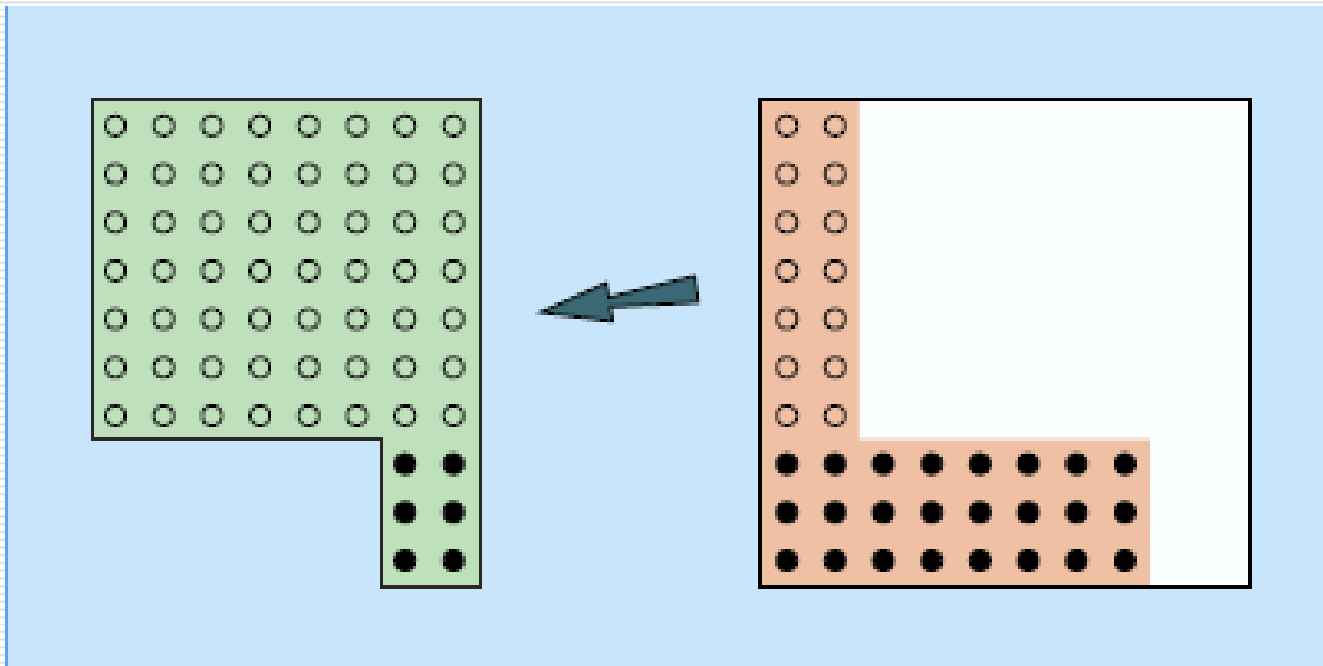
Fig 7 Perfect test

Pozitivna napovedna vrednost

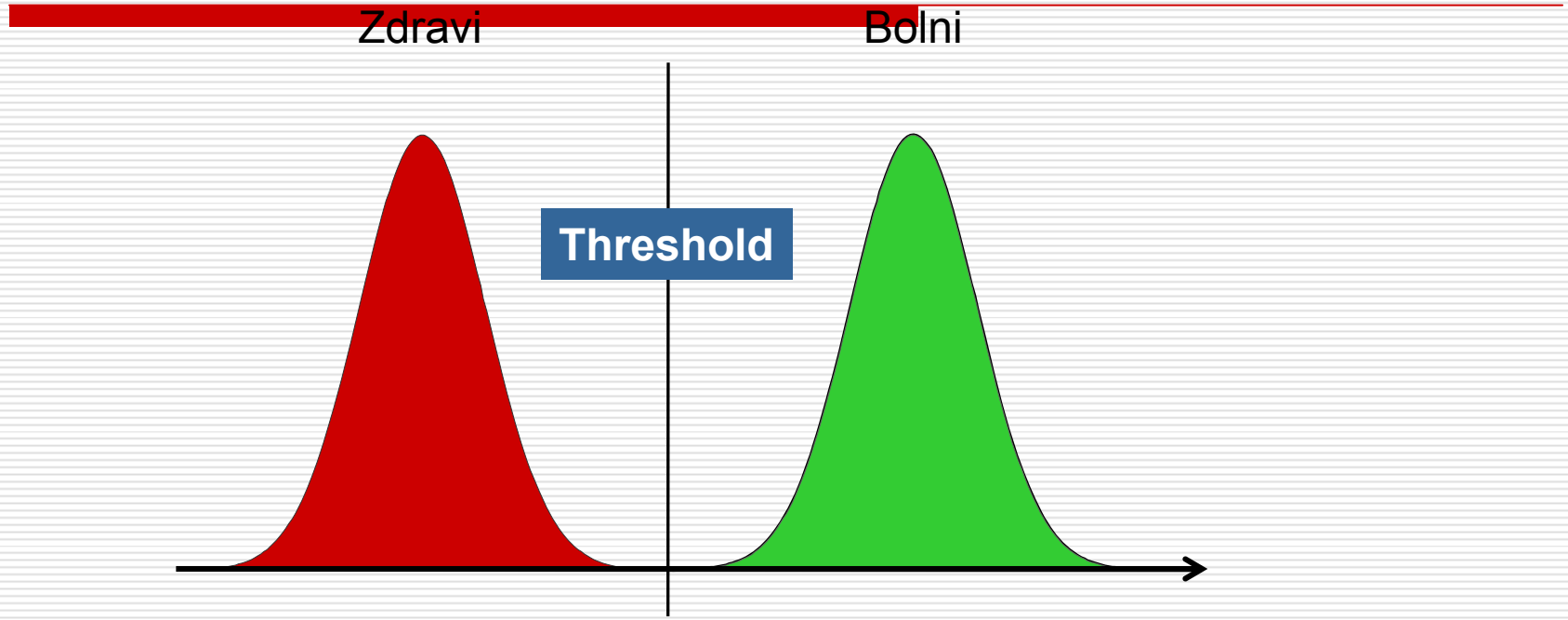


$$24/38=63\%$$

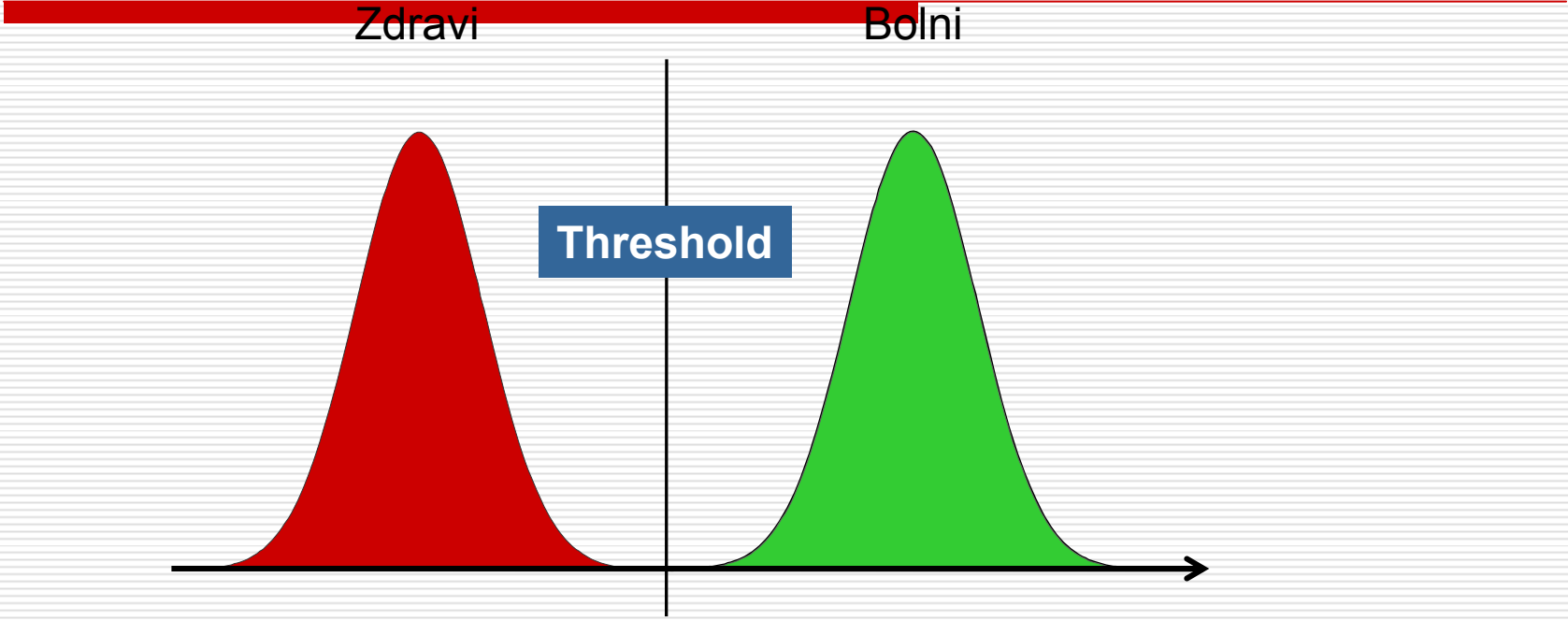
Negativna napovedna vrednost



$$56/62 = 90\%$$



Vrednost diagnostičnega parametra
ali
Subjektivna ocena verjetja bolezn

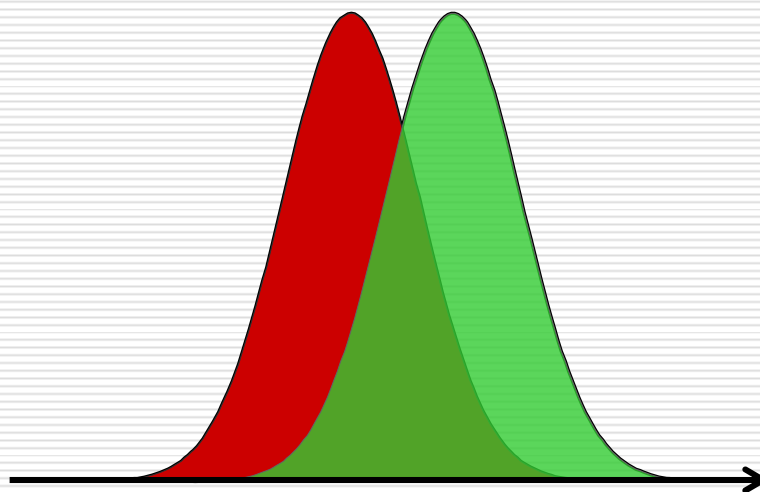


Vrednost diagnostičnega parametra
ali
Subjektivna ocena verjetja bolezni

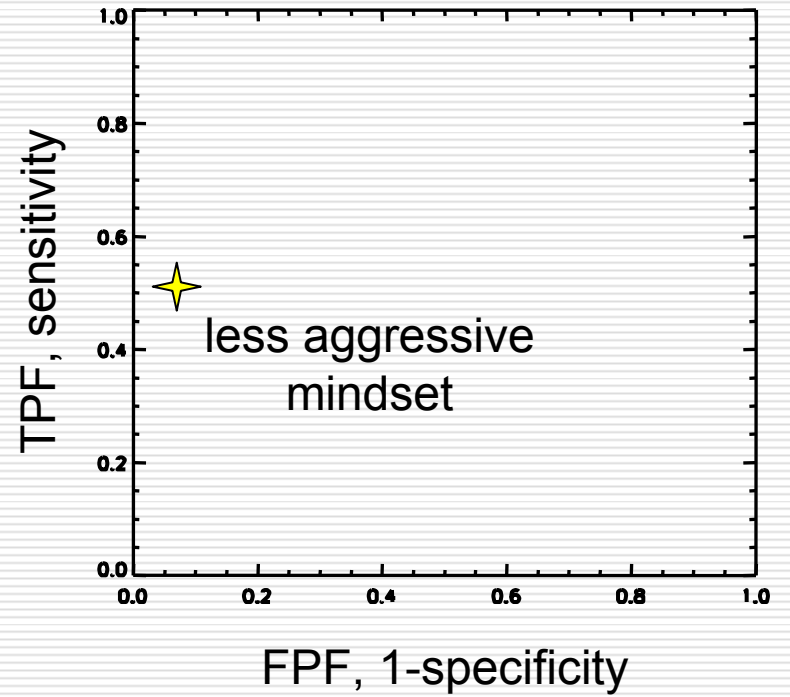
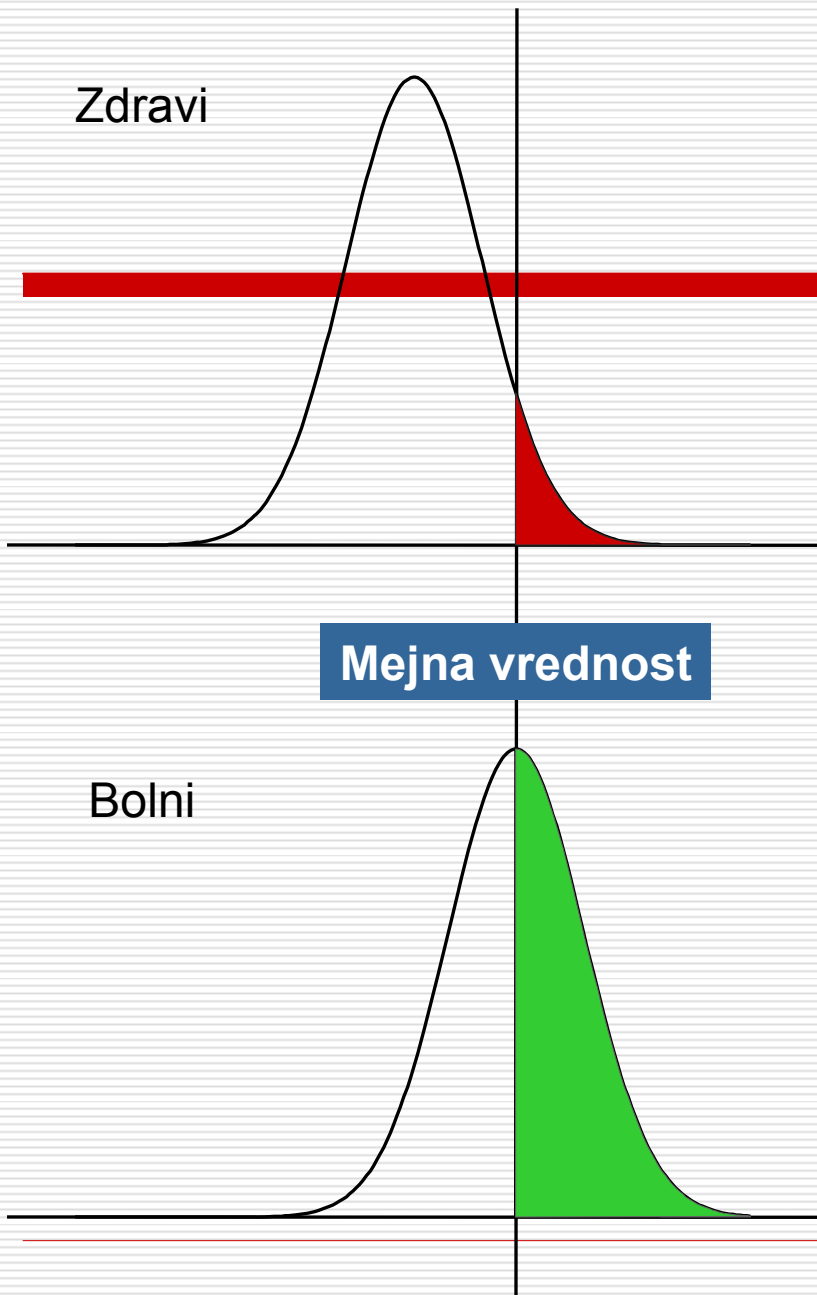
Zdravi

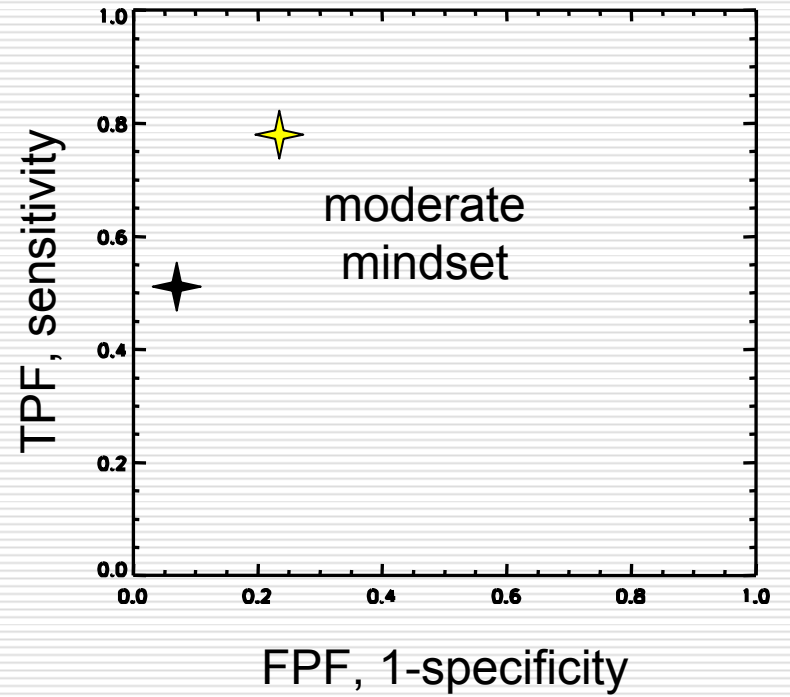
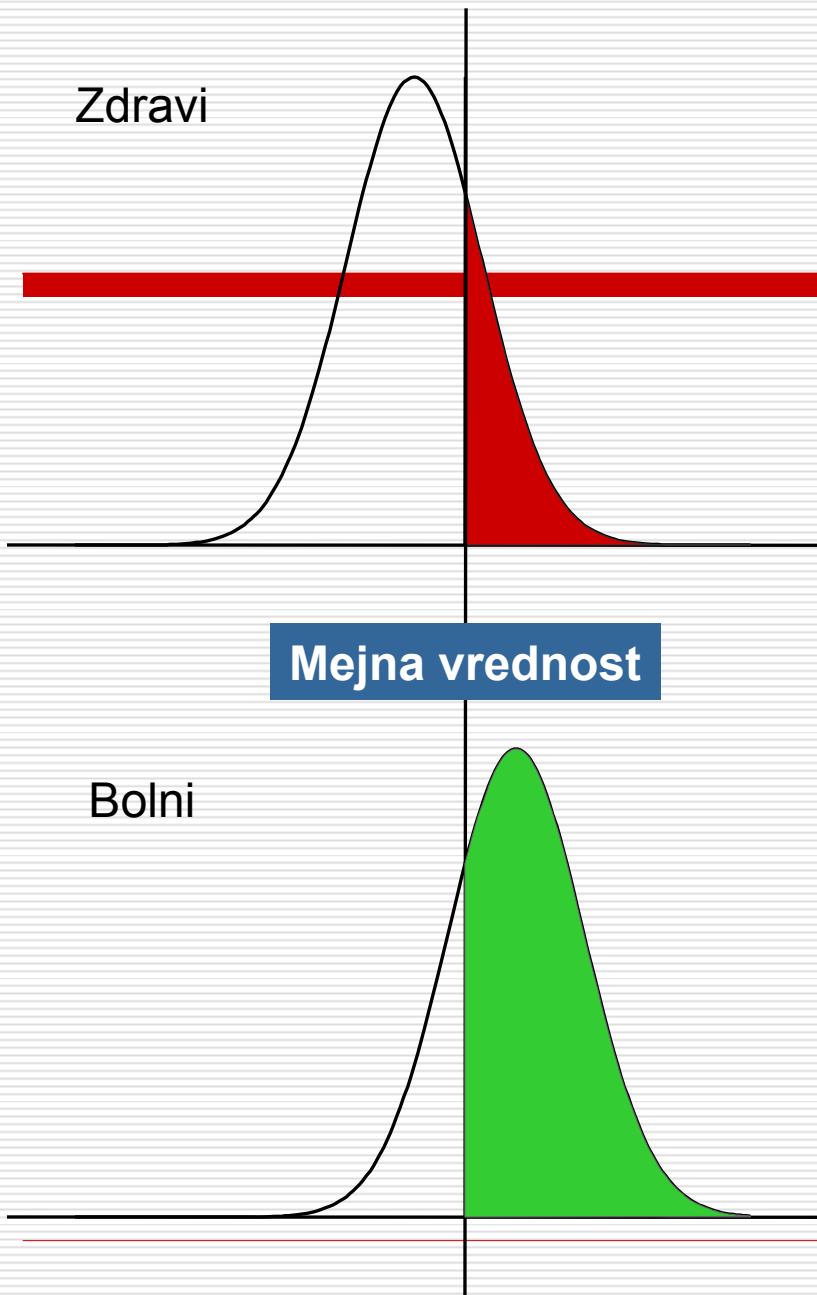
Bolni

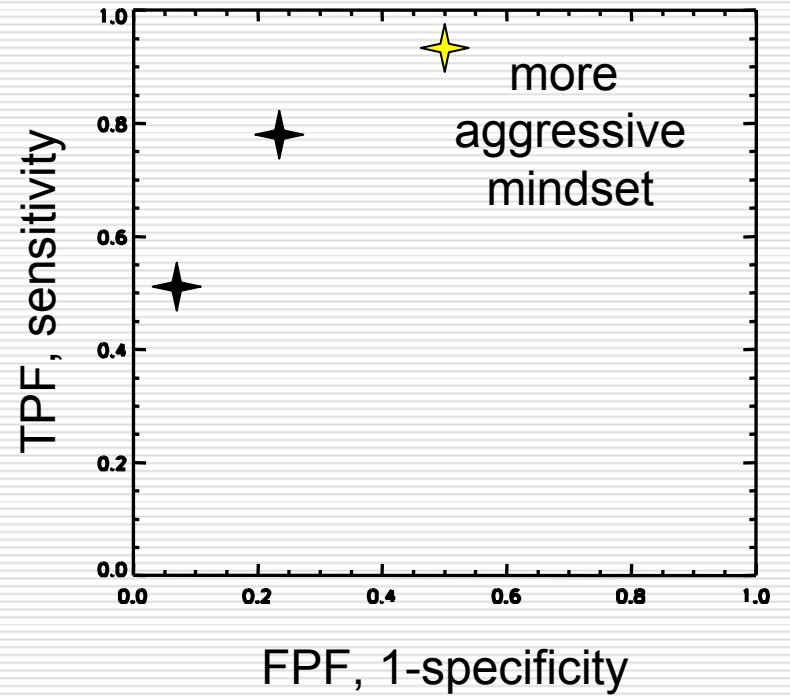
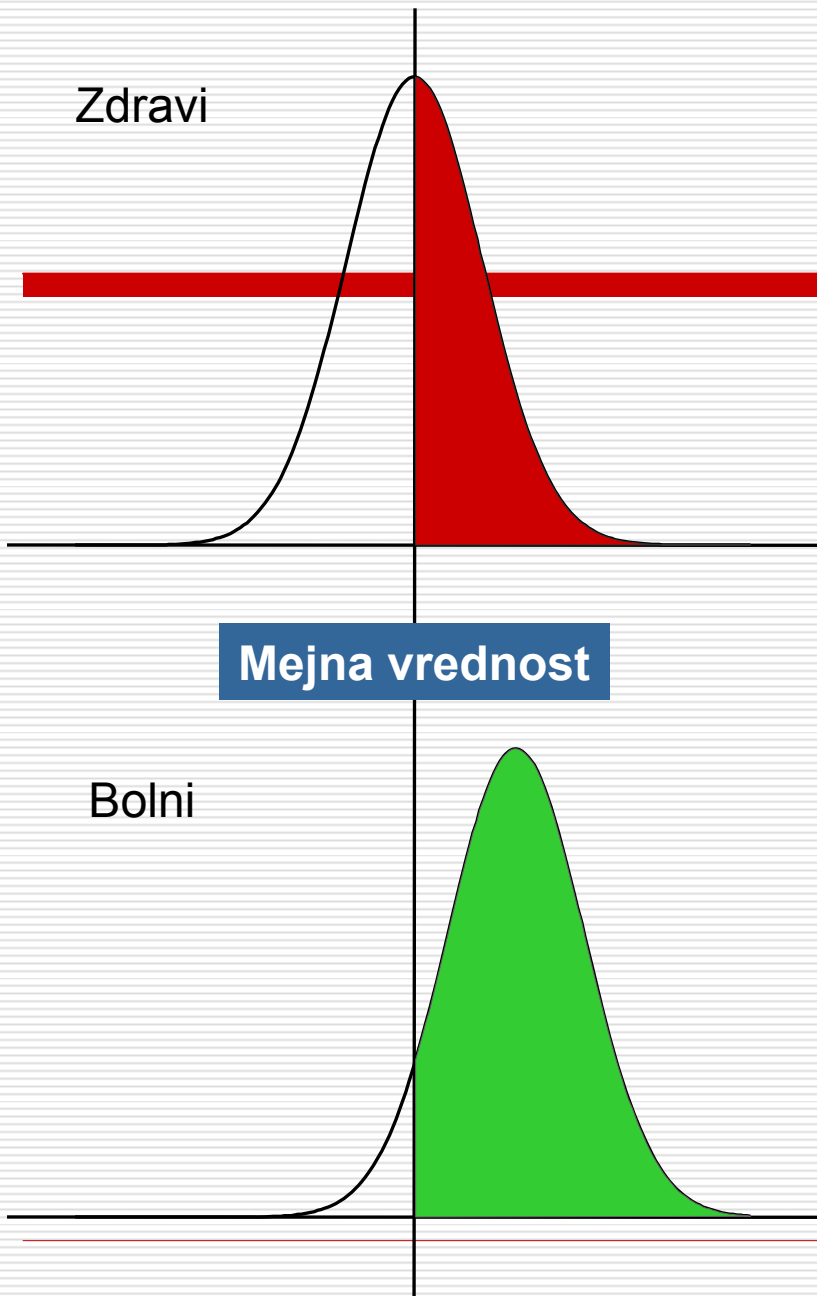
Običajneje:

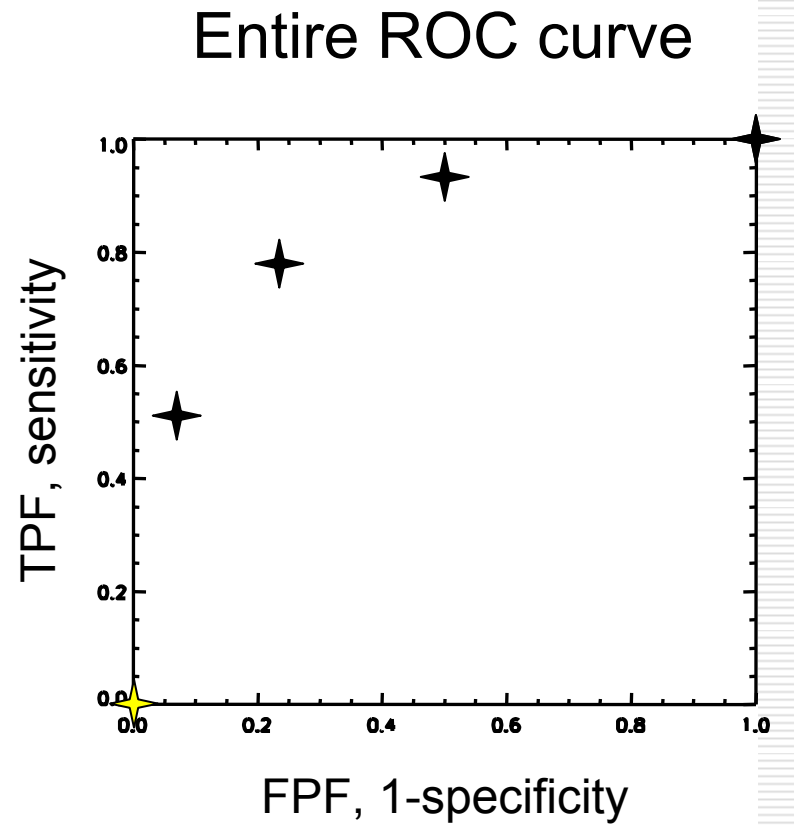
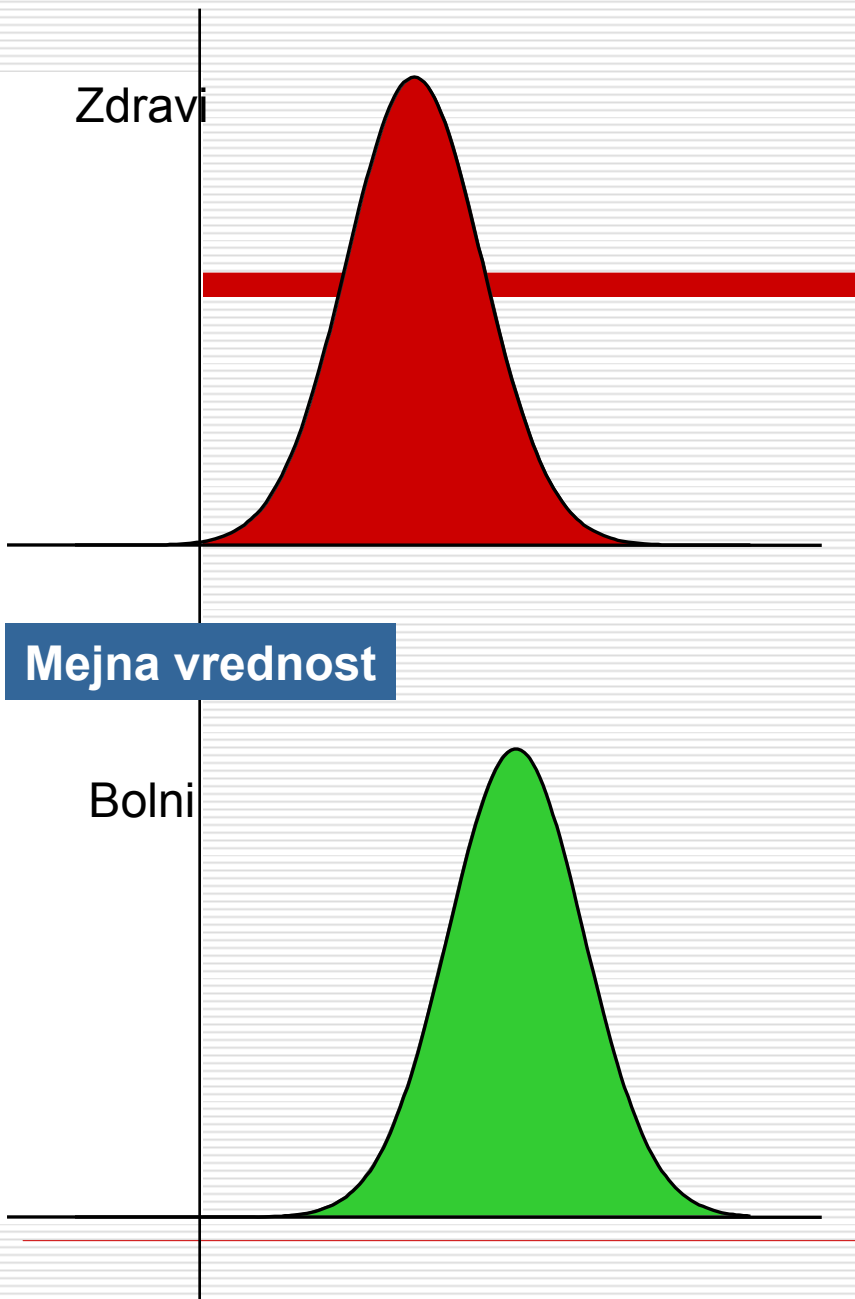


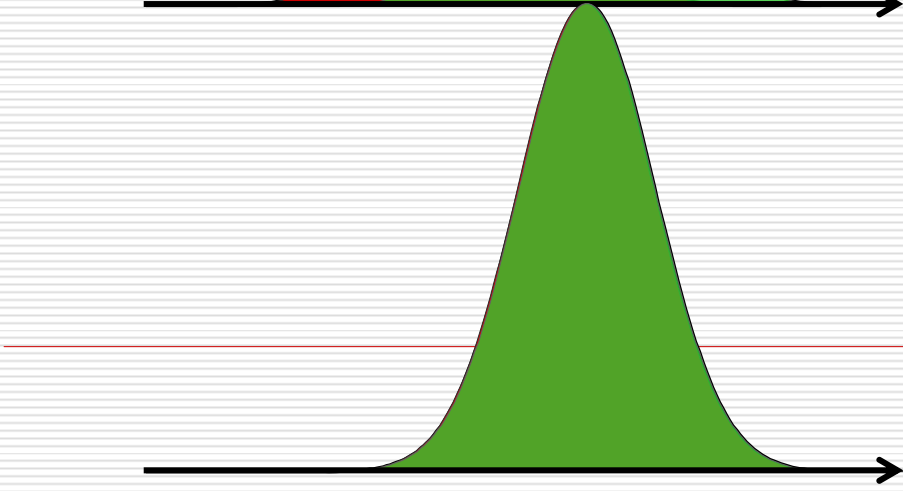
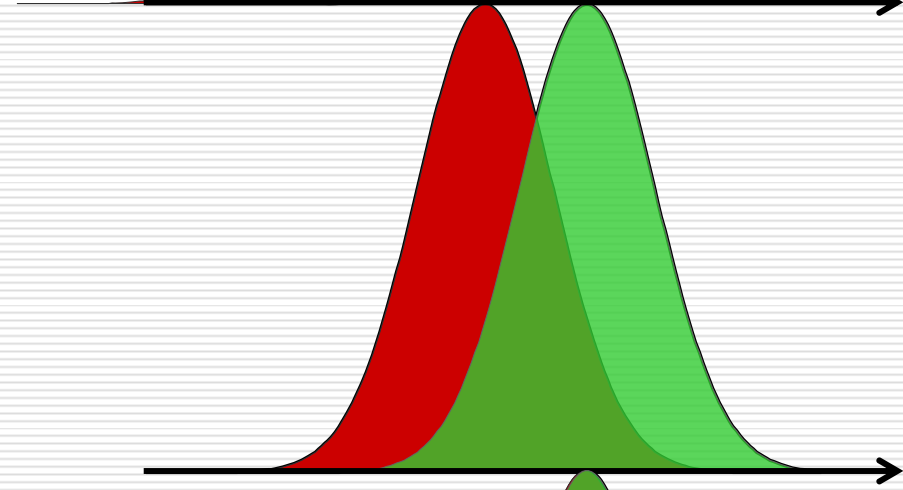
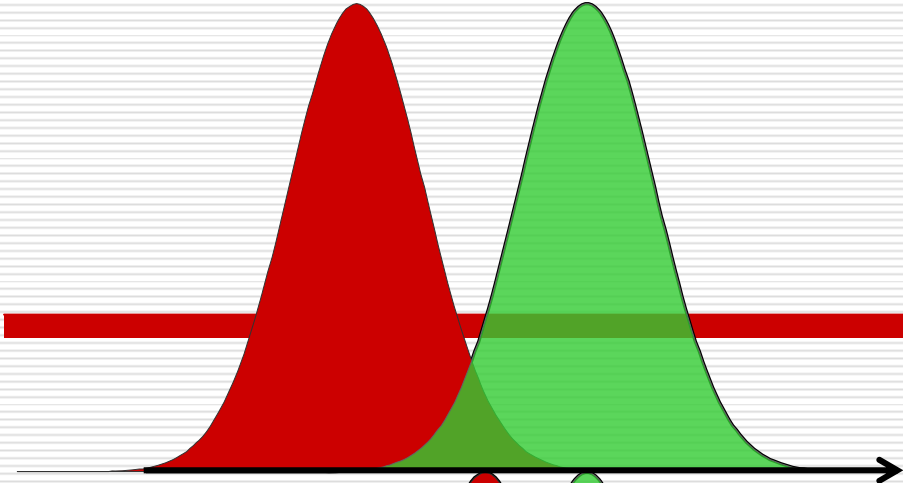
Vrednost diagnostičnega parametra
ali
Subjektivna ocena verjetja bolezn



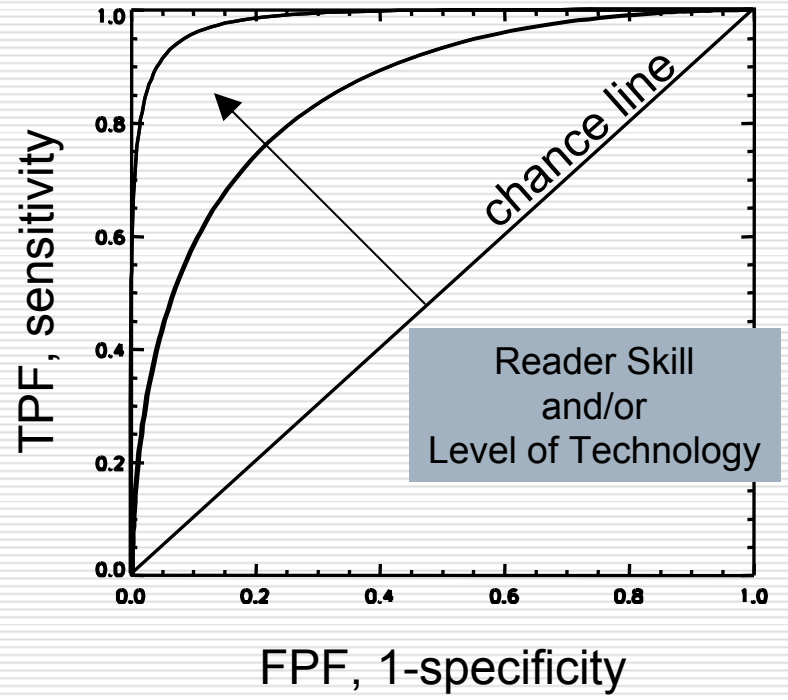








Entire ROC curve



ROC

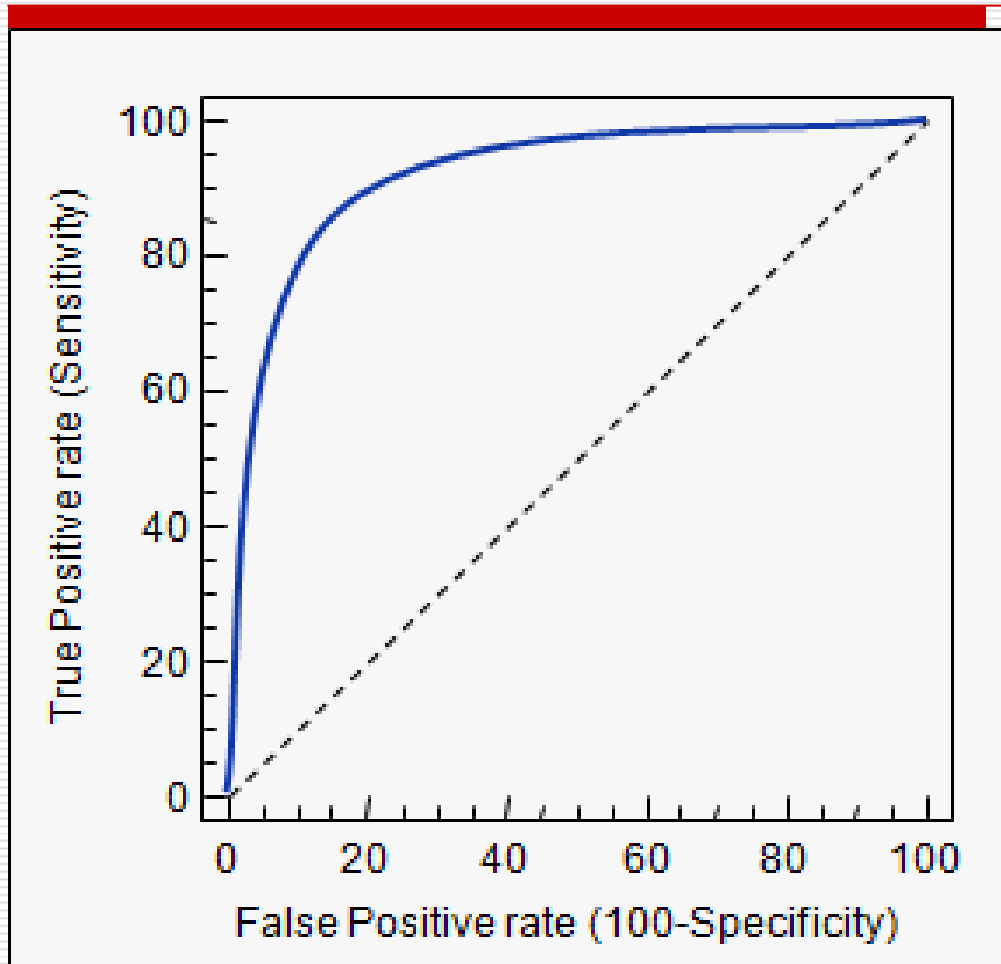
Receiver Operating Characteristic
(historično ime iz raziskav z radarjem)

Relative Operating Characteristic

Operating Characteristic

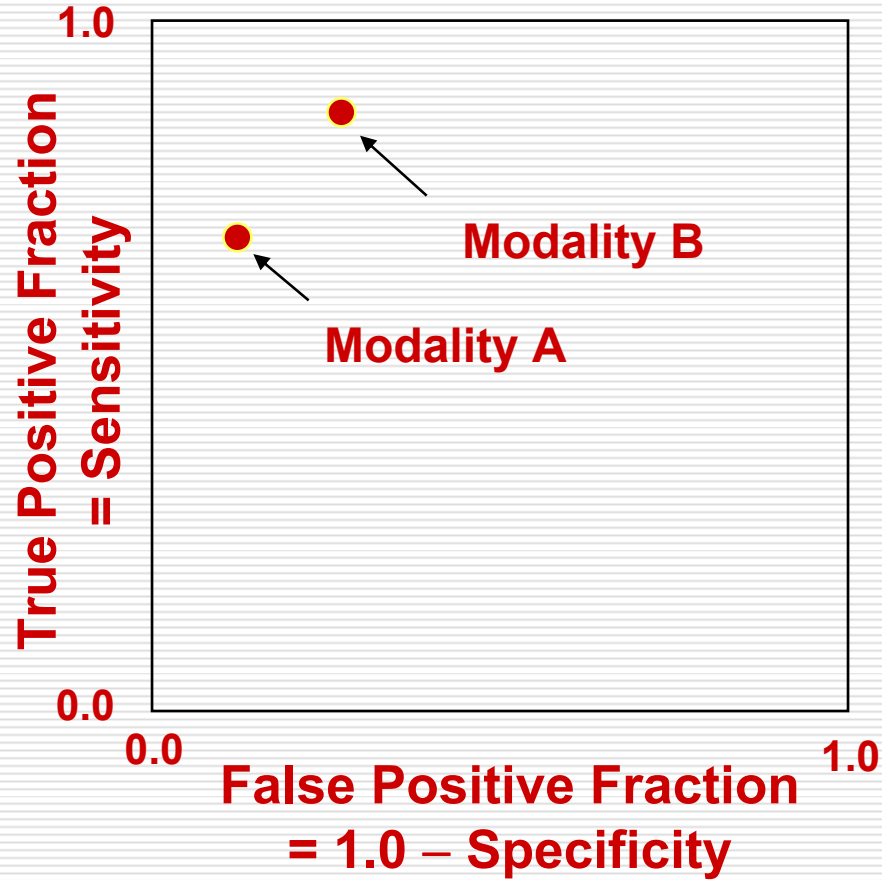
ROC

(Relative operating characteristics)

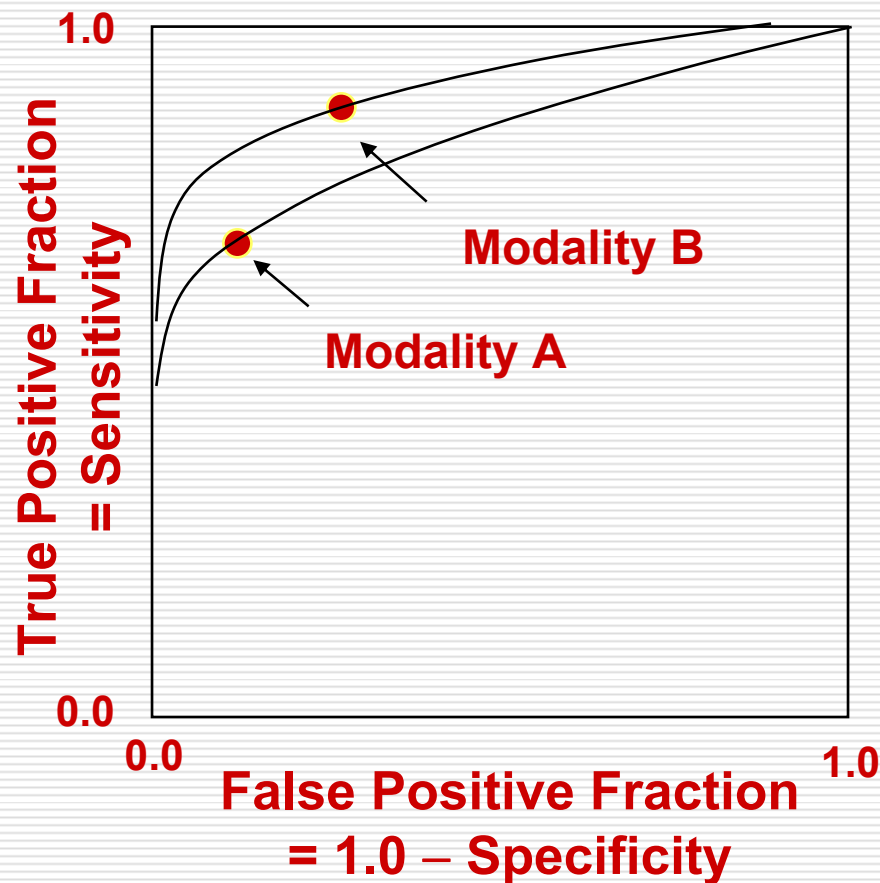


Nekaj ROC krivulj . . .

Kateri diagnostični test je boljši?



Za odgovor moramo poznati ROC krivulji (prva možnost):

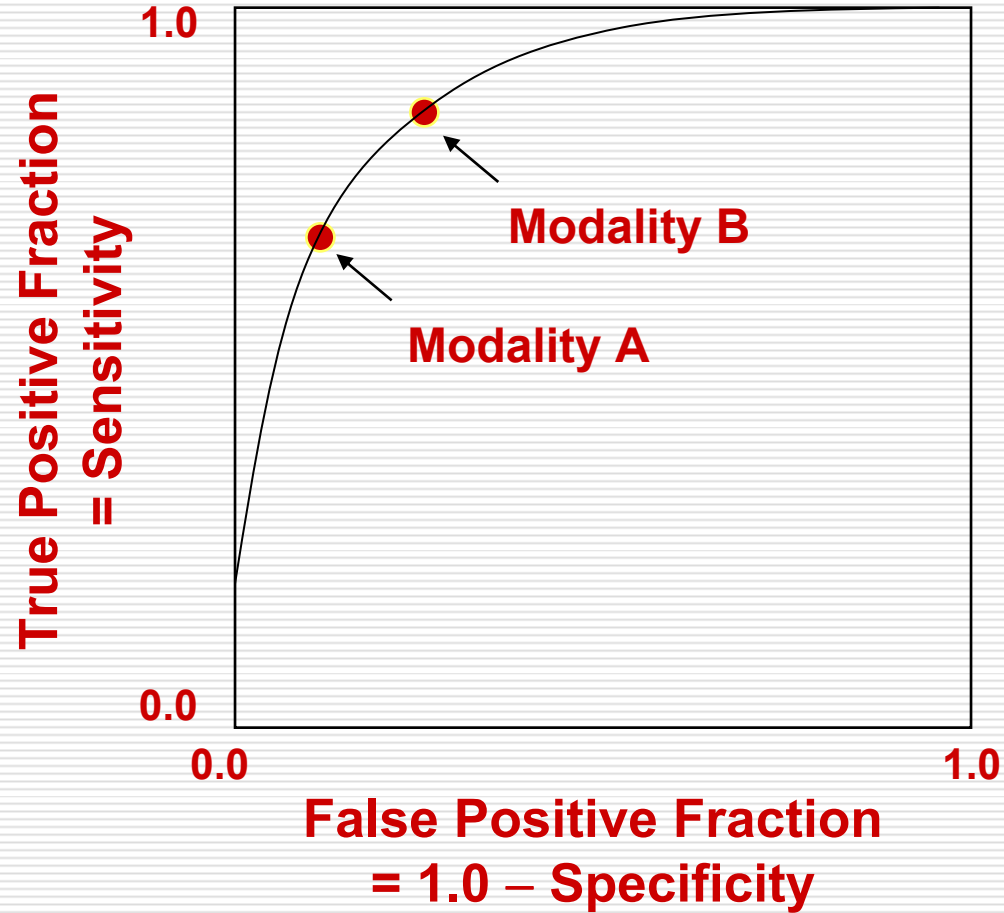


B je boljši:

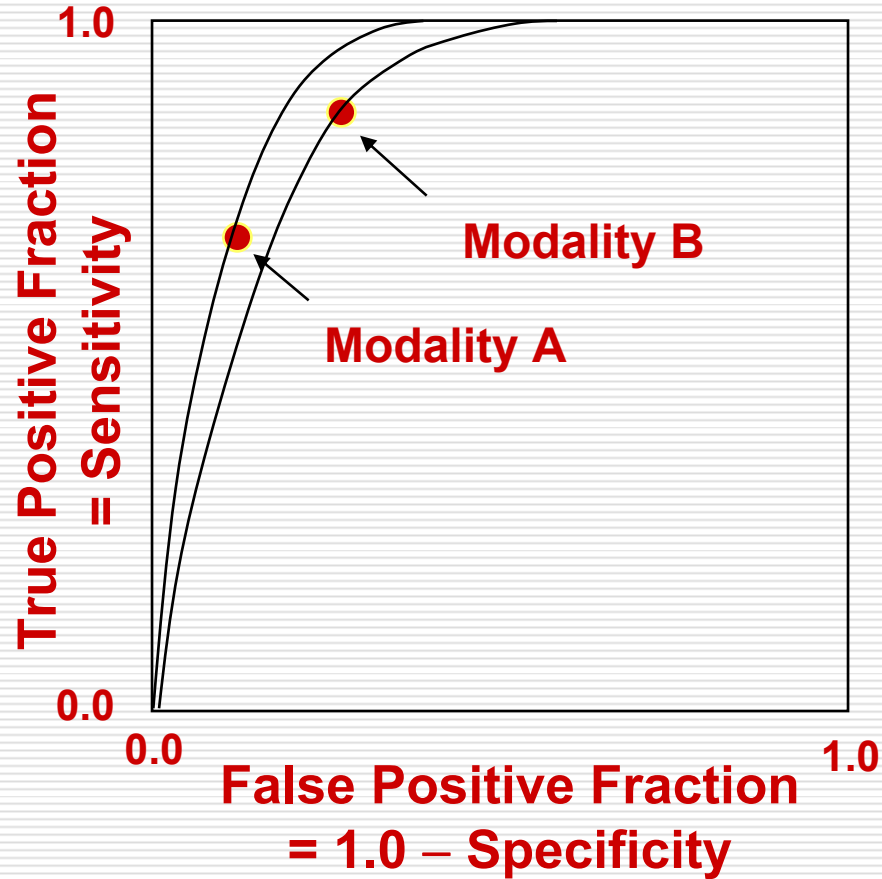
večji TPF pri
istem FPF, ali

manjši FPF pri
istem TPF

Druga možnost: Ista ROC



... In še tretja:

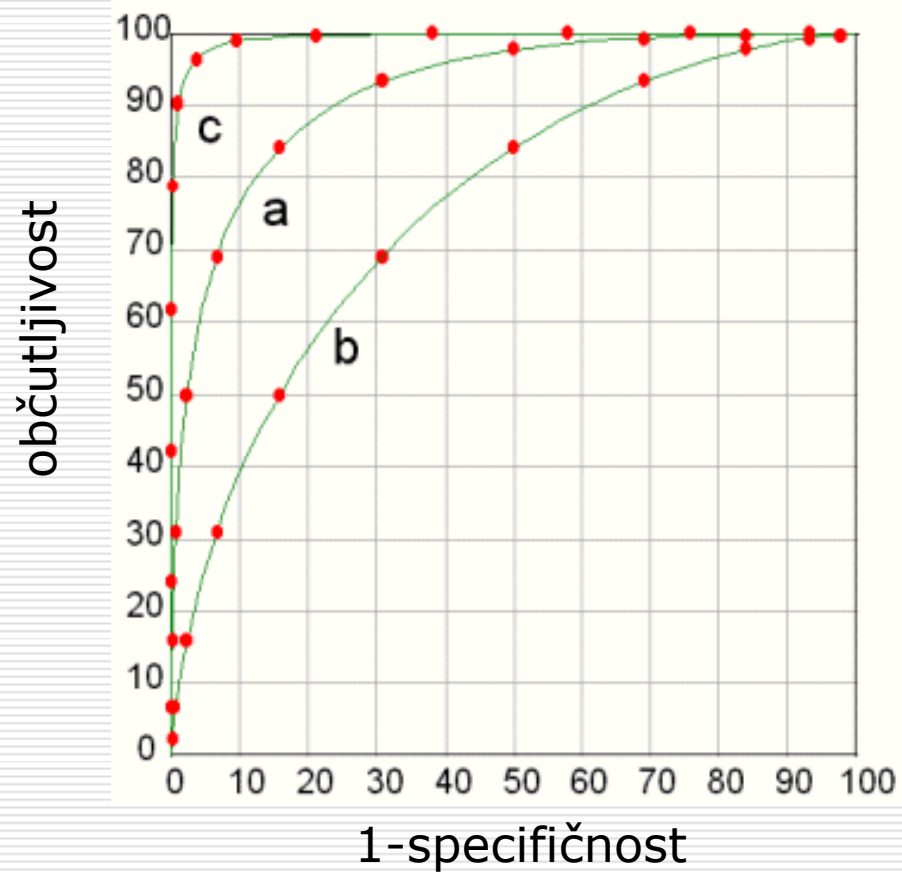


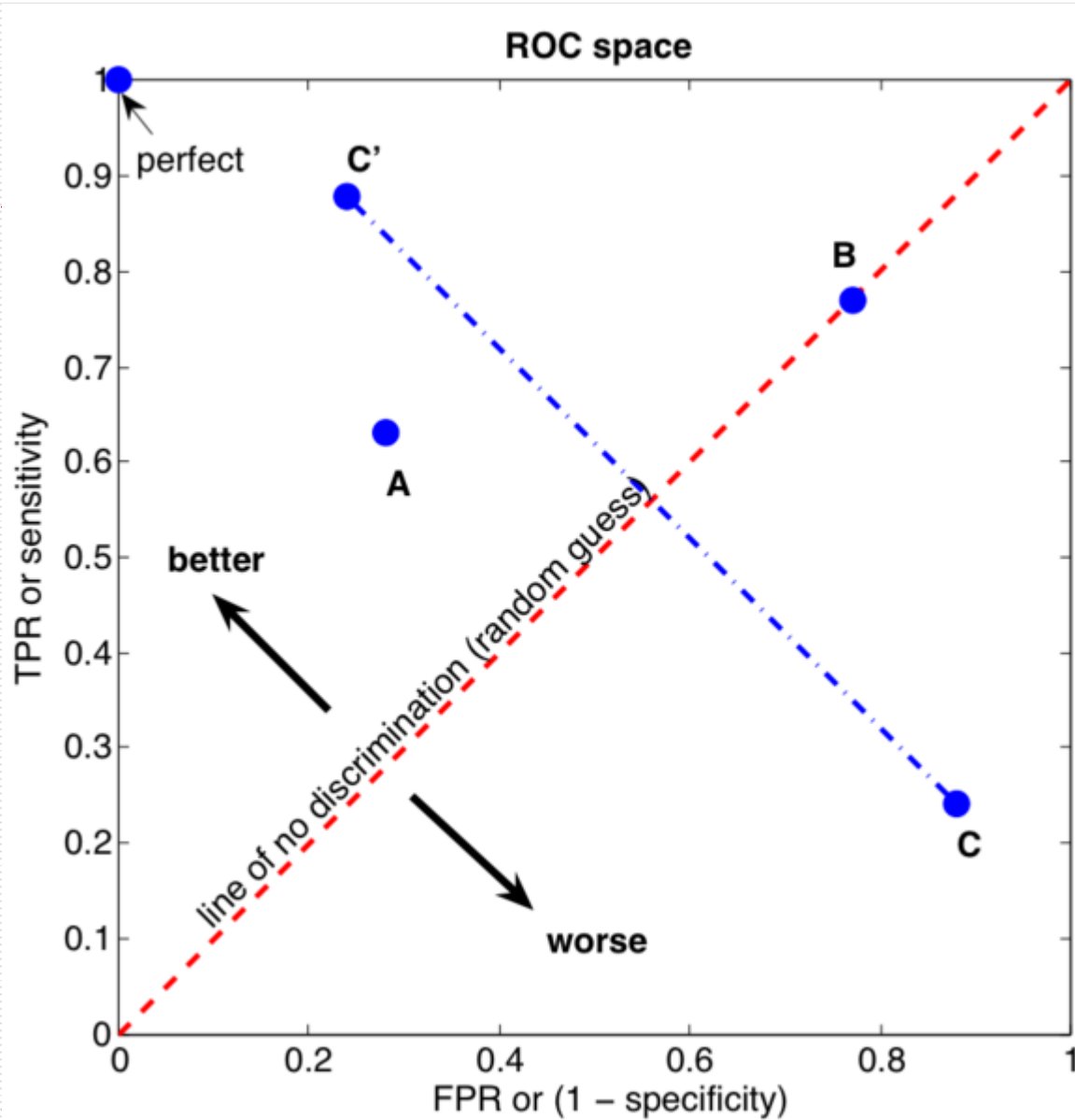
A je boljši:

večji TPF pri istem FPF, ali

manjši FPF pri istem TPF

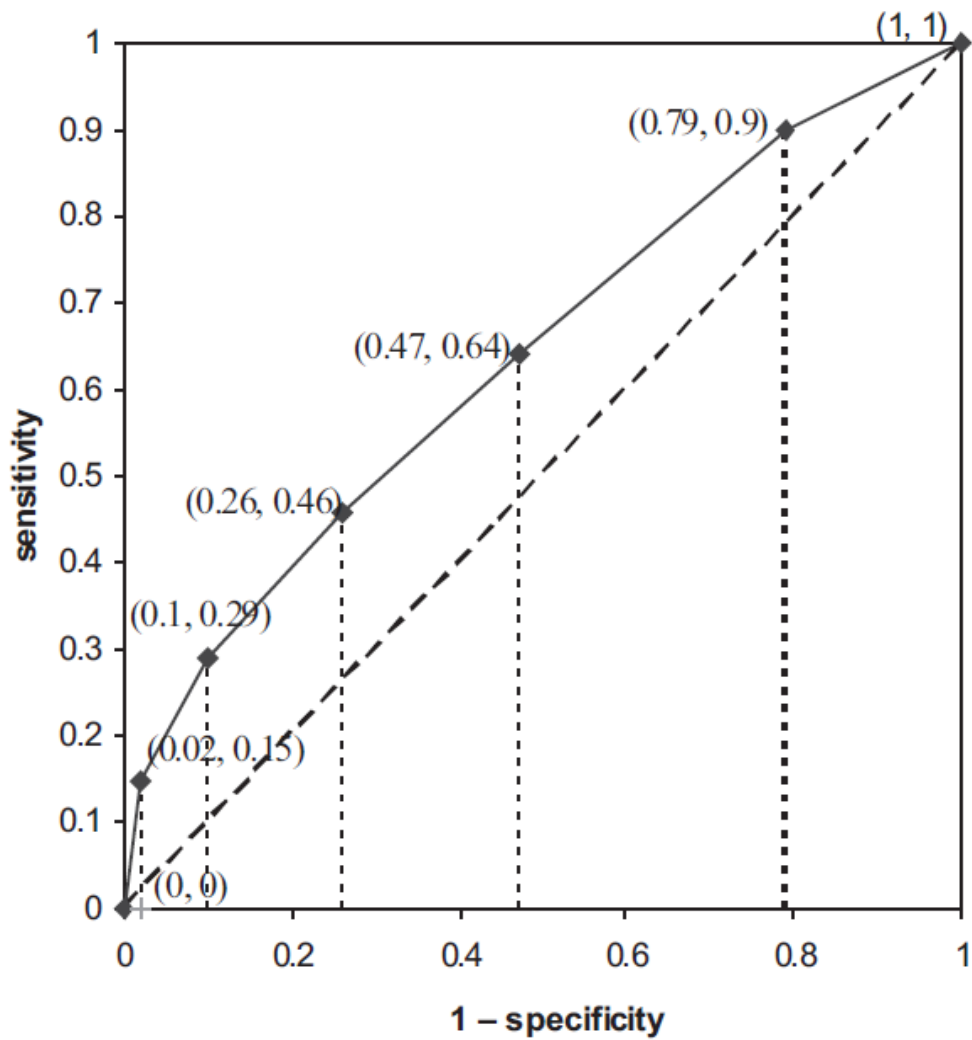
ROC





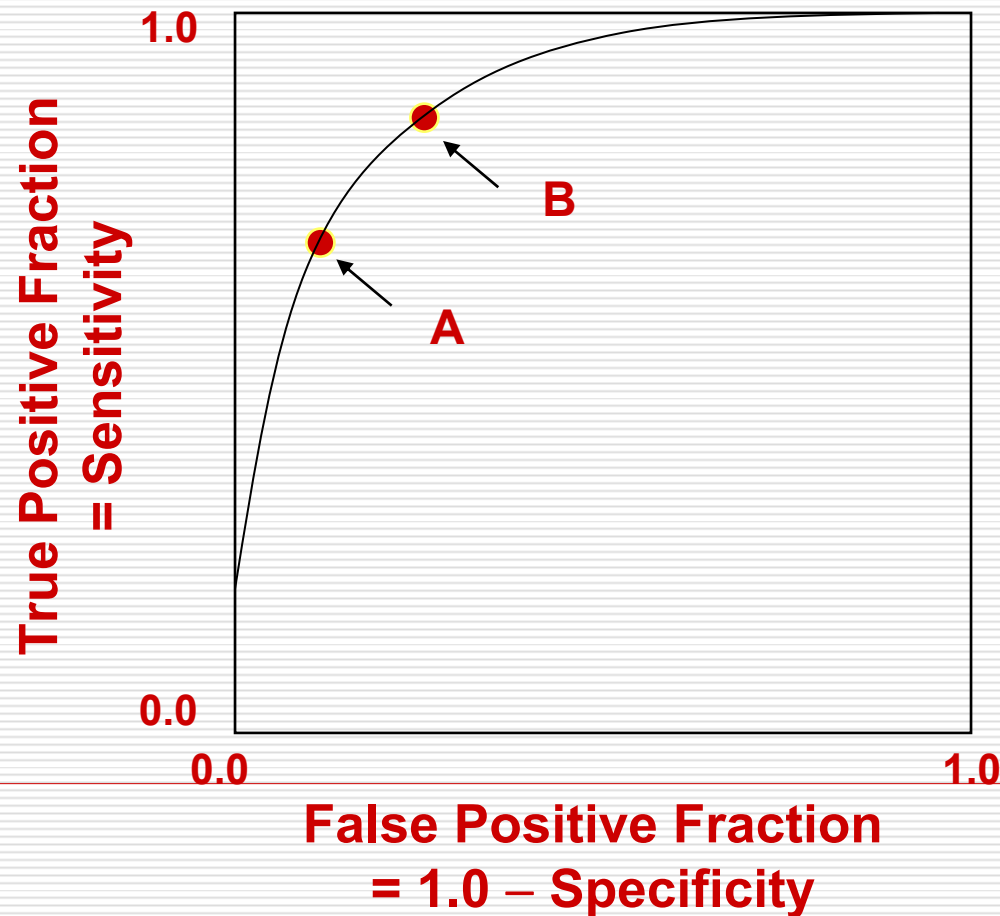
AUROC
Površina pod ROC

AUROC



Youdenov indeks

□ $J = \text{občutljivost} + \text{specifičnost} - 1$



Tiroksin in hipotiroidizem

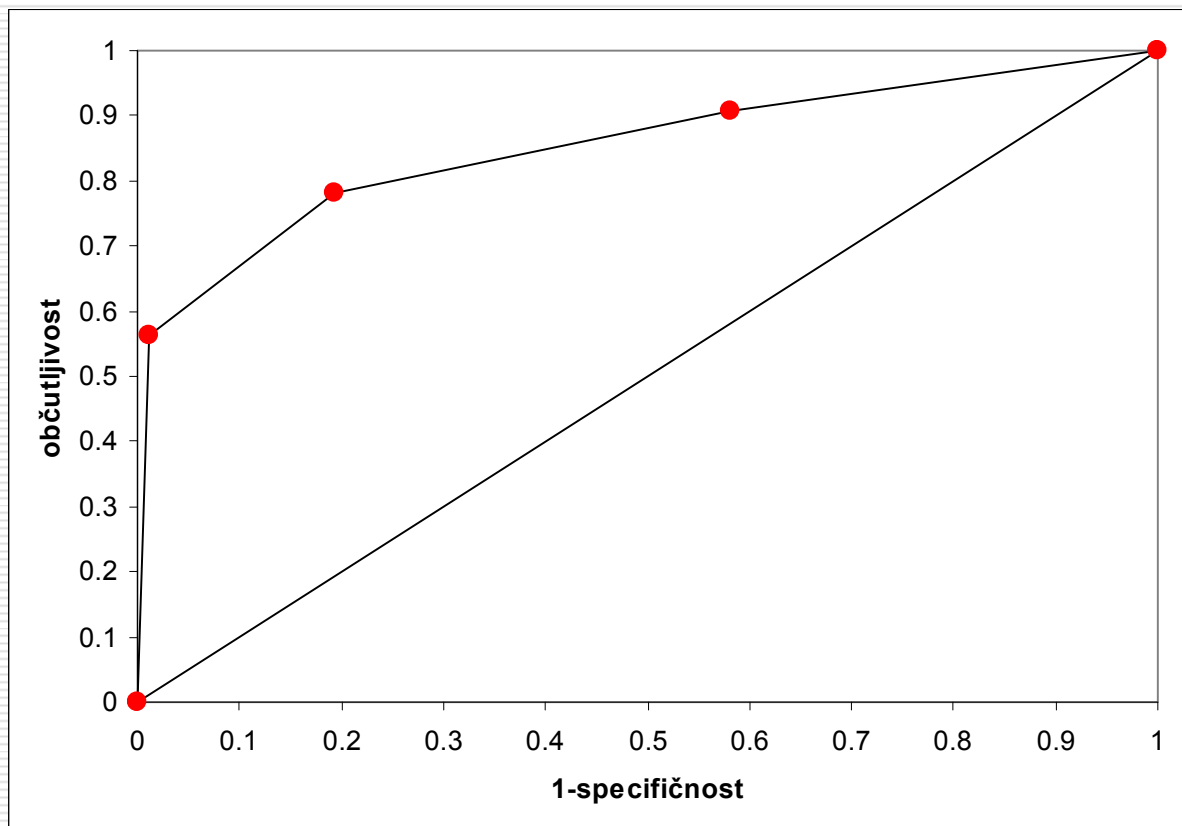
V preglednici so podani rezultati klinične diagnostike in laboratorijske analize tiroksina pri 125 bolnikih s sumom na hipotiroidizem.

Koncentracija T4 (ng/mL)	Hipotiroidizem	
	DA	NE
≤ 5	18	1
5,1 do 7,0	7	17
7,1 do 9,0	4	36
> 9	3	39

Kakšni sta specifičnost in občutljivost laboratorijskega testa za hipotiroidizem pri mejnih koncentracijah T4 5.0, 7.0 in 9.0 ng/mL?

Skiciraj ROC diagram!

ROC



THE CARTOON GUIDE TO **STATISTICS**



LARRY GONICK

*Author of **The Cartoon History of the Universe***

& WOOLLCOTT SMITH

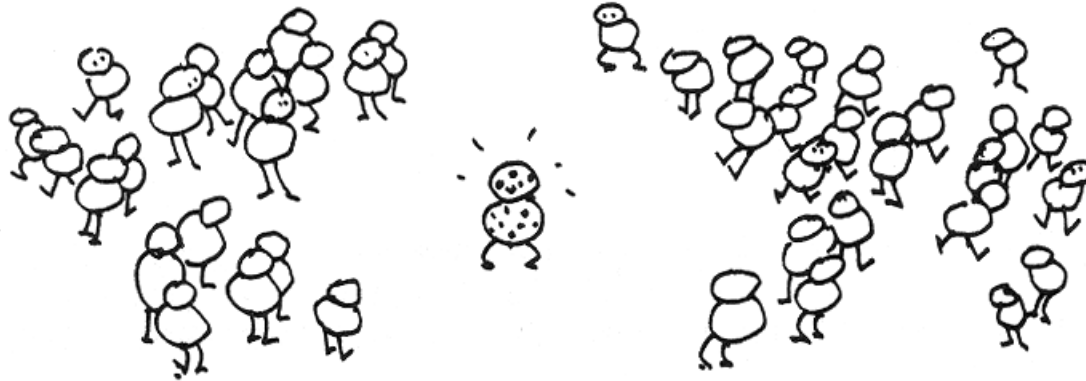
Gonick, Smith. Cartoon guide to statistics

BAYES THEOREM and the case of the false positives...

FOR A MORE SERIOUS APPLICATION OF CONDITIONAL PROBABILITY, LET'S ENTER AN ARENA OF LIFE AND DEATH...



SUPPOSE A RARE DISEASE INFECTS ONE OUT OF EVERY 1000 PEOPLE IN A POPULATION...



AND SUPPOSE THAT THERE IS A GOOD, BUT NOT PERFECT, TEST FOR THIS DISEASE: IF A PERSON HAS THE DISEASE, THE TEST COMES BACK POSITIVE 99% OF THE TIME. ON THE OTHER HAND, THE TEST ALSO PRODUCES SOME FALSE POSITIVES. ABOUT 2% OF UNINFECTED PATIENTS ALSO TEST POSITIVE. AND YOU JUST TESTED POSITIVE. WHAT ARE YOUR CHANCES OF HAVING THE DISEASE?



LET'S PUT IT THIS WAY: SHOULD I PAY IN ADVANCE?

WE HAVE TWO EVENTS TO WORK WITH:

A : PATIENT HAS THE DISEASE

B : PATIENT TESTS POSITIVE.

THE INFORMATION ABOUT THE TEST'S
EFFECTIVENESS CAN BE WRITTEN



$$P(A) = .001$$

(ONE PATIENT IN 1000 HAS THE DISEASE)

$$P(B|A) = .99$$

(PROBABILITY OF A POSITIVE TEST,
GIVEN INFECTION, IS .99)

$$P(B|NOT A) = .02$$

(PROBABILITY OF A FALSE POSITIVE, GIVEN
NO INFECTION, IS .02)

AND WE ASK

$$P(A|B) = \text{WHAT?}$$

(PROBABILITY OF HAVING THE DISEASE,
GIVEN A POSITIVE TEST)

SINCE THE TREATMENT FOR THIS DISEASE HAS SERIOUS SIDE EFFECTS, THE DOCTOR, HER LAWYER, AND HER LAWYER'S LAWYER CALL ON JOE BAYES, CP (CONSULTING PROBABILIST), FOR AN ANSWER. JOE DERIVES A THEOREM FIRST PROVED BY HIS ANCESTOR, THE REV. THOMAS BAYES (1701-1761).



I WARN YOU...
THIS IS GOING TO
USE - CACKLE -
CONDITIONAL
PROBABILITY...

JOE BEGINS WITH A 2X2 TABLE, WHICH DIVIDES THE SAMPLE SPACE INTO FOUR MUTUALLY EXCLUSIVE EVENTS. IT DISPLAYS EVERY POSSIBLE COMBINATION OF DISEASE STATE AND TEST RESULT.

	A	NOT A
B	A AND B	NOT A AND B
NOT B	A AND NOT B	NOT A AND NOT B

LET'S FIND THE PROBABILITIES OF EACH EVENT IN THE TABLE:

	A	NOT A	SUM
B	$P(A \text{ AND } B)$	$P(\text{NOT } A \text{ AND } B)$	$P(B)$
NOT B	$P(A \text{ AND NOT } B)$	$P(\text{NOT } A \text{ AND NOT } B)$	$P(\text{NOT } B)$
	$P(A)$	$P(\text{NOT } A)$	1

THE PROBABILITIES IN THE MARGINS ARE FOUND BY SUMMING ACROSS ROWS AND DOWN COLUMNS.

NOW COMPUTE:

$$P(A \text{ AND } B) = P(B|A)P(A) = (.99)(.001) = .00099$$

$$P(\text{NOT } A \text{ AND } B) = P(B|\text{NOT } A)P(\text{NOT } A) = (.02)(.999) = .01998$$

ALLOWING US TO FILL IN SOME ENTRIES:



	A	NOT A	SUM
B	.00099	.01998	.02097
NOT B	P(A AND NOT B)	P(NOT A AND NOT B)	P(NOT B)
	.001	.999	1

WE FIND THE REMAINING PROBABILITIES BY SUBTRACTING IN THE COLUMNS, THEN ADDING ACROSS THE ROWS.

THE FINAL TABLE IS:

	A	NOT A	
B	.00099	.01998	P(B)
NOT B	.00001	.97902	P(NOT B)
	.001	.999	1
	P(A)	P(NOT A)	

FROM WHICH WE DIRECTLY DERIVE

$$P(A|B) = \frac{P(A \text{ AND } B)}{P(B)} = \frac{.00099}{.02097} = .0472$$

DESPITE THE HIGH ACCURACY OF THE TEST, *LESS THAN 5%* OF THOSE WHO TEST POSITIVE ACTUALLY HAVE THE DISEASE! THIS IS CALLED THE *FALSE POSITIVE PARADOX*.

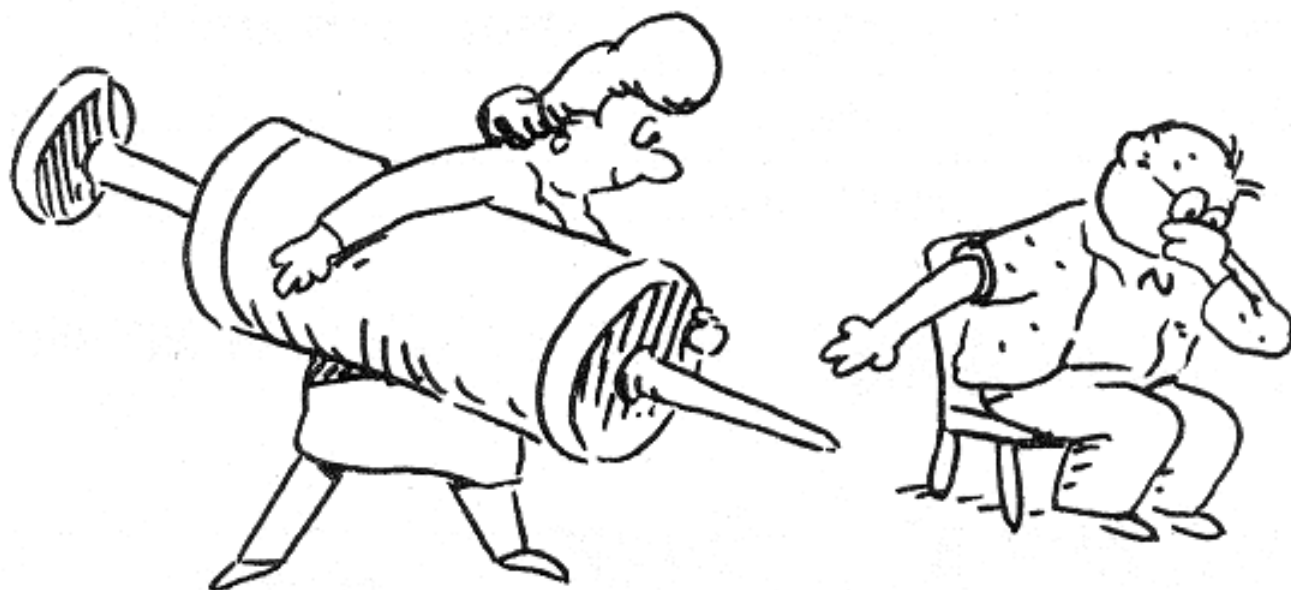
PARADOX
AND
PAIR-A-
LAWYERS...



THIS TABLE SHOWS WHAT HAPPENS IN A GROUP OF A THOUSAND PATIENTS. ON AVERAGE, ONLY 21 PEOPLE WILL TEST POSITIVE—AND ONLY *ONE* OF THEM HAS THE DISEASE! 20 FALSE POSITIVES COME FROM THE *MUCH LARGER UNINFECTED GROUP*.

	DISEASE	NO DISEASE	
TESTS POSITIVE	1	20	21
TESTS NEGATIVE	0	979	979
	1	999	1000

WHAT'S THE PHYSICIAN TO DO? JOE BAYES ADVISES HER NOT TO START TREATMENT ON THE BASIS OF THIS TEST ALONE. THE TEST DOES PROVIDE INFORMATION, HOWEVER: WITH A POSITIVE TEST THE PATIENT'S CHANCE OF HAVING THE DISEASE INCREASED FROM 1 IN 1000 TO 1 IN 23. THE DOCTOR FOLLOWS UP WITH MORE TESTS.

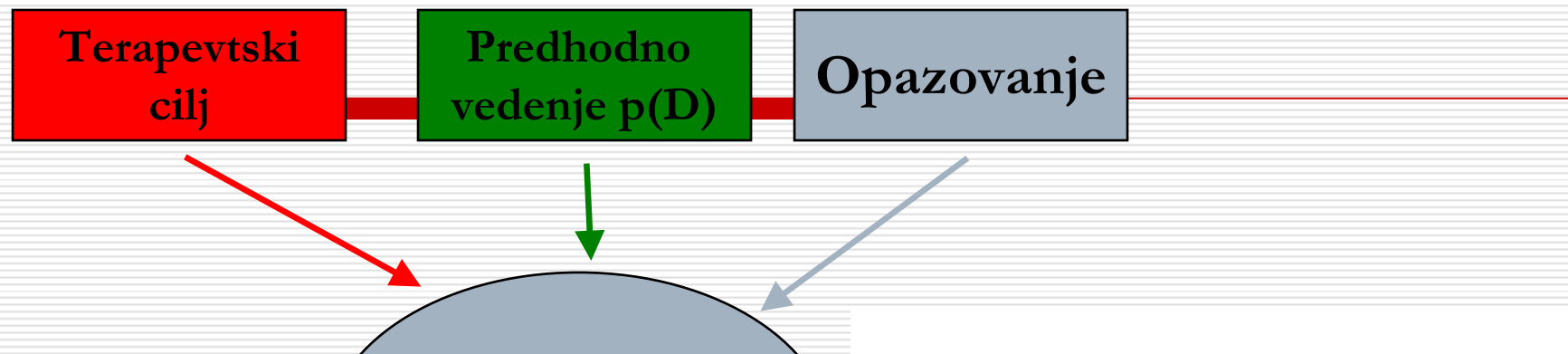


Tiroksin in hipotiroidizem

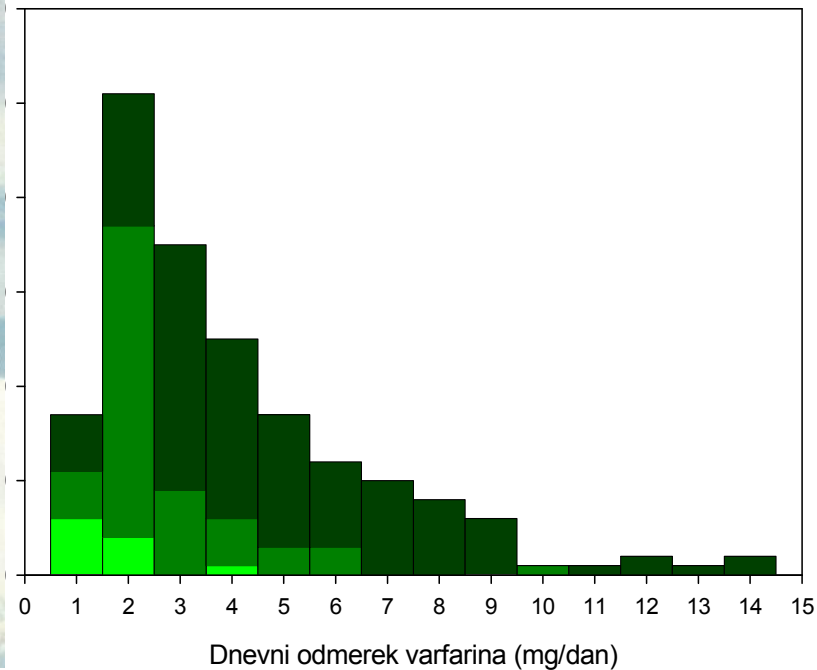
Z uporabo Bayesove formule izračunaj kakšna je pri naključno izbranem pacientu verjetnost hipotiroidizma, če je izid testa pozitiven in kot mejno vrednost izberemo 5.0, 7.0 in 9.0 ng/mL. Pri tem upoštevaj epidemiološke podatke, ki kažejo, da ima hipotiroidizem 0.25% prebivalcev?

-
- PNV se s povečevanjem prevalence povečuje.
 - Odvisna od občutljivosti in specifičnosti diagnostičnega testa.
-

Bayesovo adaptivno vodenje



odmerjanja



Thomas Bayes (1702-1761)

Primer

Kri v blatu pri bolnikih z rakom debelega črevesja

Rezultat endoskopije

		+	-	
Kri v blatu	+	20	180	200
	-	10	1820	1830
		30	2000	2030

Občutljivost, specifičnost, PNV, NNV
