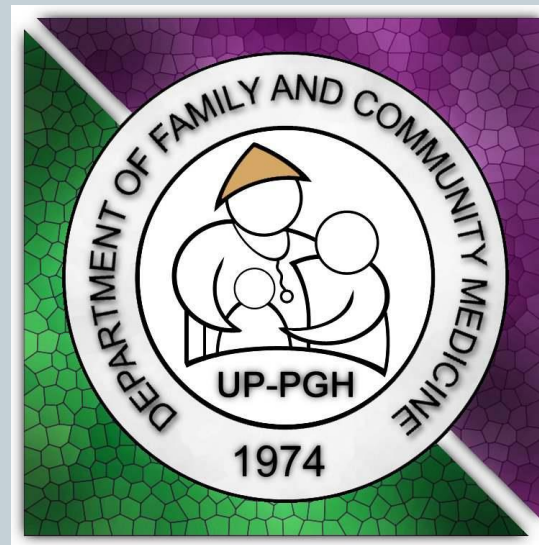


Introduction to EVIDENCE-BASED MEDICINE



DEPARTMENT OF FAMILY & COMMUNITY MEDICINE
UP-PHILIPPINE GENERAL HOSPITAL



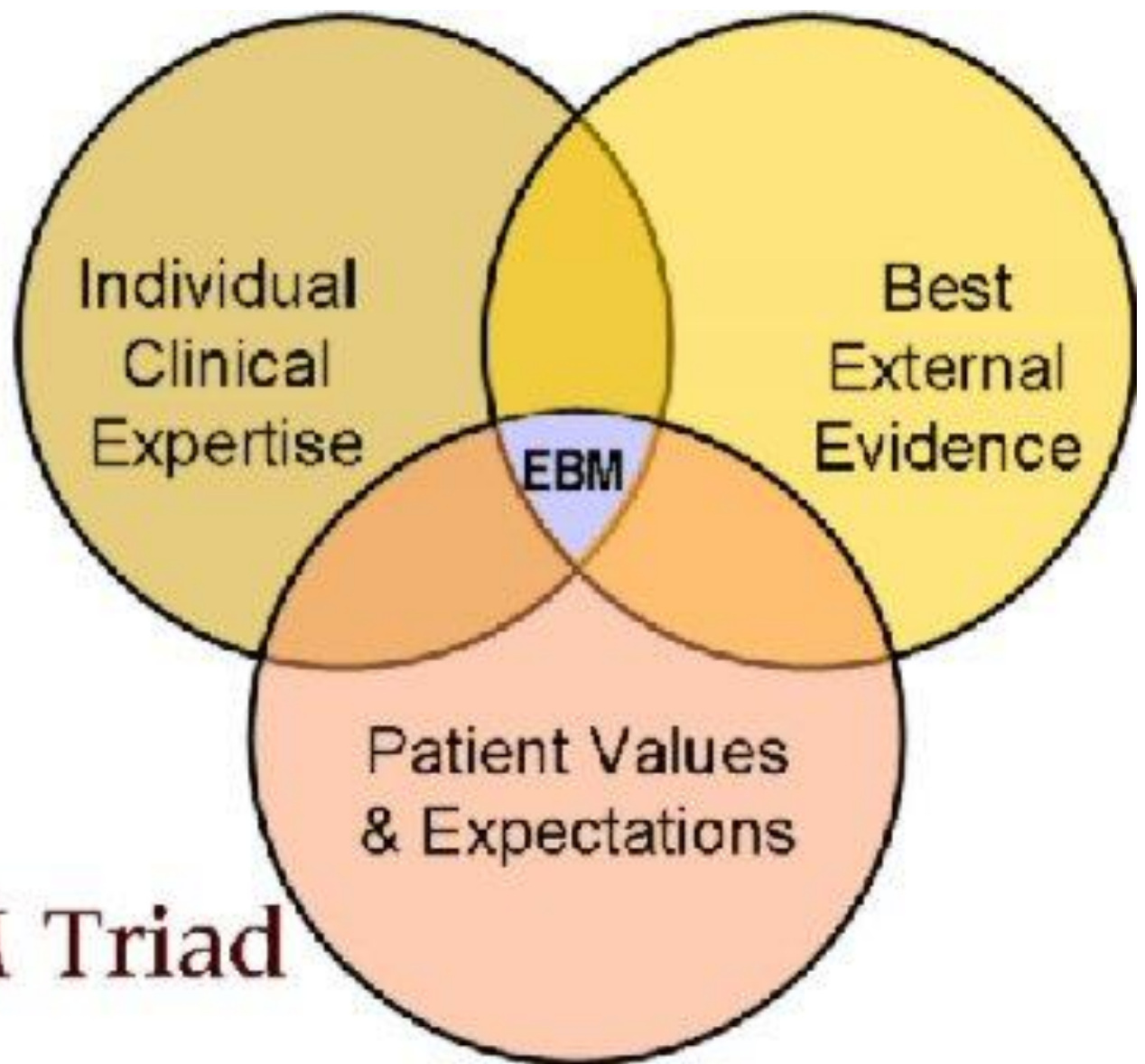
WHAT IS EVIDENCE BASED MEDICINE?



Dr. David Sackett, 1996

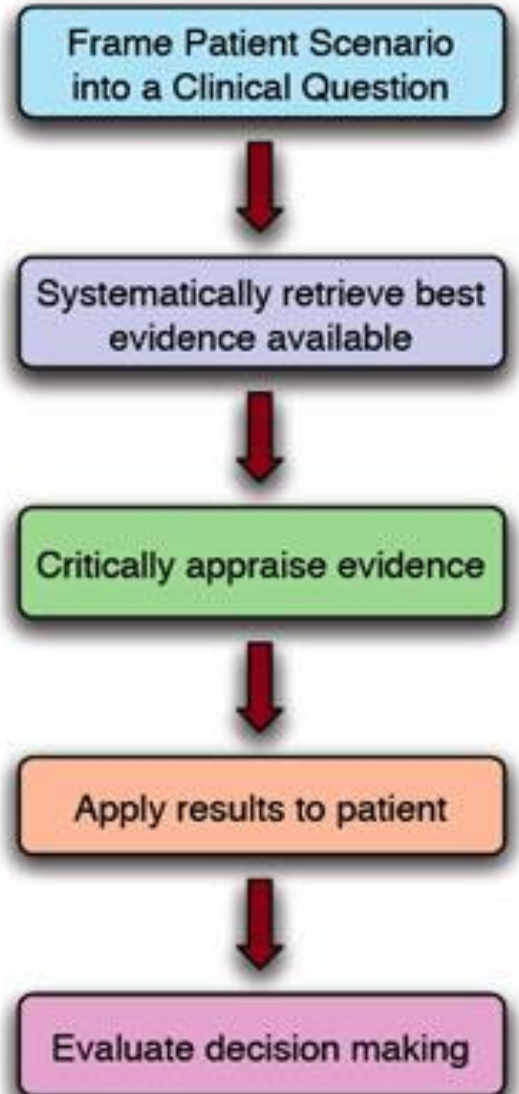
The conscientious, explicit, judicious use of current best evidence in making decisions about the care of individual patient.

It means integrating individual clinical expertise with the best available external clinical evidence from systematic research.



The EBM Triad

The Five Steps of EBM



EVIDENCE-BASED MEDICINE

STEPS IN EBM PROCESS



The patient	1. Start with the patient – a clinical problem or question arises out of the care of the patient
The question	2. Construct a well built clinical question from the case
The resource	3. Select the appropriate resource(s) and conduct the search
The evaluation	4. Appraise the evidence for its validity and applicability
The patient	5. Return to the patient – integrate the evidence with the clinical expertise, patient preference and apply it to practice
Self evaluation	6. Evaluate your performance with this patient

STEP 1: CLINICAL SCENARIO



- Maria, a 67 year old
- History of congestive heart failure brought on by several myocardial infarctions.
- Hospitalized 2x in the last 6 months for worsening of heart failure



STEP 1: CLINICAL SCENARIO



- Presently in normal sinus rhythm
- She is extremely compliant about taking her medications (enalapril, aspirin and simvastatin)
- She *desperately* wants to stay out of the hospital
- You think she should also be taking digoxin, but **you are not certain** if this will help keep her out of the hospital.
- You decide to research this question before her next visit.

STEP 2. CONSTRUCT A WELL BUILT CLINICAL QUESTION

ANATOMY OF A CLINICAL QUESTION

POPULATION	<ul style="list-style-type: none">• primary problem, disease, or co-existing conditions• sex, age or race of a patient
INTERVENTION OR TREATMENT	<ul style="list-style-type: none">• main intervention, prognostic factor, or exposure• prescribe a drug, Order a test? Order surgery?
COMPARISON OR CONTROL	<ul style="list-style-type: none">• main alternative to compare with the intervention• Another drug? Another test? Placebo?• Your clinical question does not always need a specific comparison
OUTCOME	<ul style="list-style-type: none">• What do you hope to accomplish/measure/improve/affect?• What are you trying to do for the patient?• Symptoms? Number of adverse events? Functionality? Test scores?
METHODOLOGY	<ul style="list-style-type: none">• Type of Evidence/Study

GOING BACK TO MARIA



Patient / Problem	Elderly with congestive heart failure
Intervention	Digoxin
Comparison	Placebo
Outcome	Primary: reduce number of hospitalizations Secondary: reduce mortality
Methodology	Randomized Controlled Trial



TYPE OF QUESTION: WHAT TYPE OF STUDY?



Type of Question	Suggested best type of Study
Therapy	RCT>cohort > case control > case series
Diagnosis	Cross sectional>prospective, blind comparison to a gold standard
Etiology/Harm	RCT > cohort > case control > case series
Prognosis	cohort study > case control > case series
Prevention	RCT>cohort study > case control > case series
Cost	economic analysis

What type of study is best suited for Maria's case?

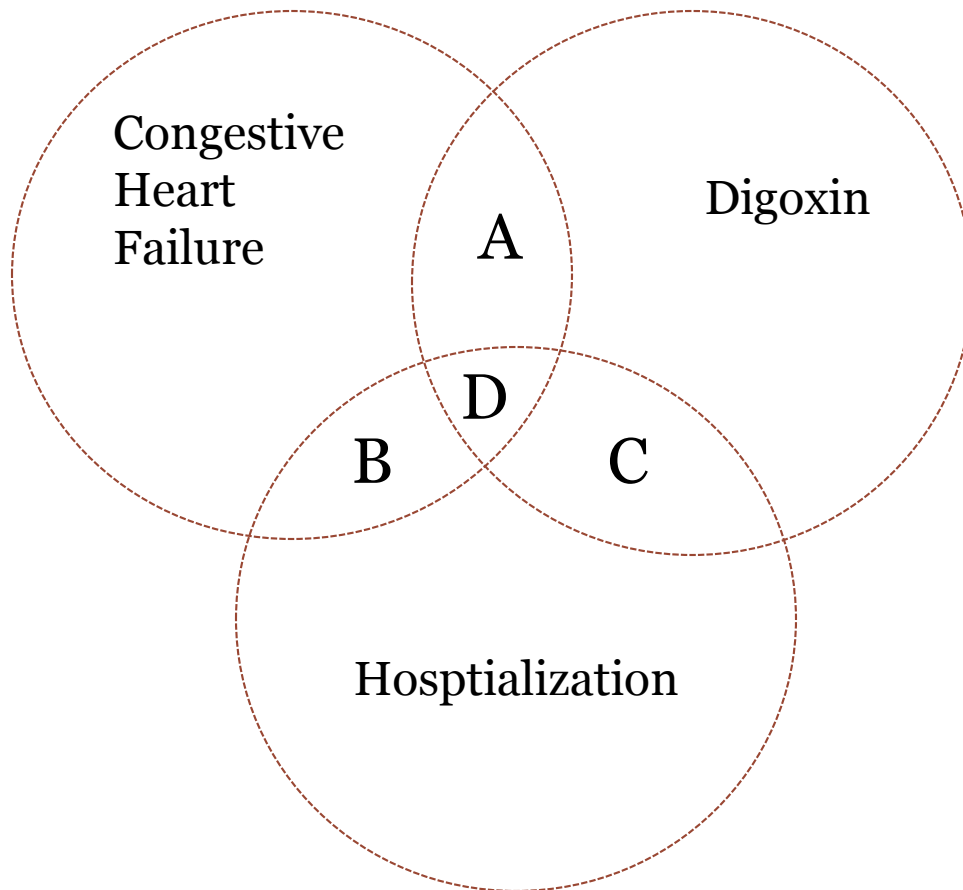
STEP 3: SELECT THE APPROPRIATE RESOURCE AND CONDUCT THE SEARCH



SEARCH STRATEGIES

- Identify the concepts: **KEY TERMS**
- Phrase search: “quotation marks”
- Boolean Principle: **OR, AND**
- MeSH
- Truncation and Wild Card
- Limits
- Related Articles
- Clinical Queries

BOOLEAN PRINCIPLE



By using OR &
AND...

How do we get

- A?
- B?
- C?
- D?

Filters: [Manage Filters](#)

New feature

Results by year

PMC Images search for "congestive heart failure"

Titles with your search terms

2382 free full-text articles in PubMed Central

Find related data

Search details

Recent Activity

Clear all

[Show additional filters](#)

Article types

- Clinical Trial
- Review
- More ...

Text availability

- Abstract available
- Free full text available
- Full text available

PubMed Commons

- Reader comments

Publication dates

- 5 years
- 10 years
- Custom range...

Species

- Humans
- Other Animals

Clear all

Show additional filters

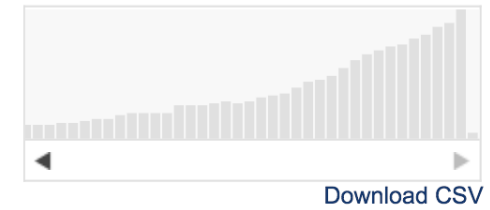
Results: 1 to 20 of 168573

- ☐ [2-\[18F\]-fluoro-2-desoxy-D-glucose positron emission tomography initial staging impacts on survival in Hodgkin lymphoma.](#)
 Cerci JJ, Linardi CC, Pracchia LF, Junior JS, Trindade E, Delbeke D, Cerci RJ, Carr R, Meneghetti JC, Buccheri V.
 World J Radiol. 2013 Dec 28;5(12):484-90. doi: 10.4329/wjr.v5.i12.484.
 PMID: 24379935 [PubMed]
[Related citations](#)
- ☐ [A theory-based exercise intervention in patients with heart failure: A protocol for randomized, controlled trial.](#)
 Rajati F, Mostafavi F, Sharifirad G, Sadeghi M, Tavakol K, Feizi A, Pashaei T.
 J Res Med Sci. 2013 Aug;18(8):659-67.
 PMID: 24379841 [PubMed]
[Related citations](#)
- ☐ [Short-term vs. long-term heart rate variability in ischemic cardiomyopathy risk stratification.](#)
 Voss A, Schroeder R, Vallverdú M, Schulz S, Cygankiewicz I, Vázquez R, Bayés de Luna A, Caminal P.
 Front Physiol. 2013 Dec 13;4:364. doi: 10.3389/fphys.2013.00364.
 PMID: 24379785 [PubMed]
[Related citations](#)
- ☐ [Proteinuria and its relation to cardiovascular disease.](#)
 Currie G, Delles C.

New feature

Try the new Display Settings option - Sort by Relevance


Results by year



Related searches

- congestive heart failure treatment
- congestive heart failure readmission
- congestive heart failure review
- acute congestive heart failure
- congestive heart failure pathophysiology

PMC Images search for congestive heart failure

Display Settings:  Summary, 20 per page, Sorted by Recently Added

Send to: 

Filters: [Manage Filters](#)

Results: 1 to 20 of 78

<< First < Prev Page 1 of 4 Next > Last >>

Text availability

Full text available

Publication dates

Custom range...

Species

Other Animals

Clear all

[Show additional filters](#)

[Digoxin: the results of the DIG study in the XXI century].

1. Helber I, Tucci PJ.

Arq Bras Cardiol. 2010 Oct;95(4):e108-11. Multiple languages.

PMID: 21180781 [PubMed - indexed for MEDLINE] [Free Article](#)

Related citations

☐ BNP-guided vs symptom-guided heart failure therapy: the Trial of Intensified vs
2. Standard Medical Therapy in Elderly Patients With **Congestive Heart Failure**
(TIME-CHF) randomized trial.

Pfisterer M, Buser P, Rickli H, Gutmann M, Erne P, Rickenbacher P, Vuillomenet A, Jeker U, Dubach P, Beer H, Yoon SI, Suter T, Osterhues HH, Schieber MM, Hilti P, Schindler R, Brunner-La Rocca HP; TIME-CHF Investigators.

JAMA. 2009 Jan 28;301(4):383-92. doi: 10.1001/jama.2009.2.

PMID: 19176440 [PubMed - indexed for MEDLINE]

Related citations

- ☐ Atrial fibrillation and congestive heart failure.

3. Roy D, Talajic M, Dubuc M, Thibault B, Guerra P, Macle L, Khairiy P.

Curr Opin Cardiol. 2009 Jan;24(1):29-34. Review.

PMID: 19102036 [PubMed - indexed for MEDLINE]

Related citations

- Drug treatment of chronic heart failure in the elderly.

4. Leibundgut G, Pfisterer M, Brunner-La Rocca HP.

Drugs Aging. 2007;24(12):991-1006. Review.

New feature

Try the new Display Settings option - Sort by Relevance

2 free full-text articles in PubMed Central

Enhanced External Counterpulsation
(E[Ont Health Technol Assess Ser. 2006]

Atrial fibrillation after cardiac surgery: a major morbid event? [Ann Surg. 1997]

[See all \(2\)...](#)

Find related data

Database:

Search details

```
"congestive heart failure"  
[All Fields] AND ("digoxin"  
[MeSH Terms] OR "digoxin"  
[All Fields]) AND  
("hospitalisation"[All
```

[Digoxin: the results of the X

Warning: Something's Not X

← → ↺

www.ncbi.nlm.nih.gov/pubmed/21180781

☆ 🔊 ☰

NCBI

Resources ▾ How To ▾

Sign in to NCBI

PubMed.gov

US National Library of Medicine
National Institutes of Health

PubMed ▾

Advanced

Search

Help

Display Settings: ▾ Abstract

Send to: ▾ free full text available at SciELO.org

Arq Bras Cardiol. 2010 Oct;95(4):e108-11.

[Digoxin: the results of the DIG study in the XXI century].

[Article in Multiple languages]

Helber I, Tucci PJ.

Author information

Abstract

After the report that there was no statistical significance in the general mortality of the DIG study, the indication of digoxin in the treatment regimens for congestive heart failure (CHF) drastically decreased. Post hoc studies that reassessed the DIG study data, indicated that an aspect that was not considered in this multicenter study has a critical influence on the prognosis of patients: the serum levels of digoxin. Regarding those that received a placebo, the general mortality and hospitalization were decreased in patients with a digoxin level < 0.9 ng/ml. At the first study that assessed the influence of digitalis in an experimental model of CHF, we verified in our lab that female rats with congestive syndrome secondary to myocardial infarction have a prolonged survival when undergoing treatment with digitoxin. The current information recommends that the merits of digoxin continue to be analyzed in order to adequately establish its importance in the treatment of CHF.

PMID: 21180781 [PubMed - indexed for MEDLINE] Free full text

Publication Types, MeSH Terms, Substances

LinkOut - more resources

PubMed Commons

PubMed Commons home

0 comments

Save items

☆ Add to Favorites ▾

Related citations in PubMed

Association of serum digoxin concentration and outcome [JAMA. 2003]

Review Digoxin remains useful in the management [Med Clin North Am. 2003]

[Effect of digoxin on atrial natriuretic peptide (ANI) [Pol Arch Med Wewn. 2001]

Review Digoxin use in congestive heart failure. Current status. [Drugs. 1998]

Review [The digitalis: should we forget it?]. [G Ital Cardiol (Rome). 2008]

See reviews...

See all...

Related information

Related Citations

Compound (MeSH Keyword)

MedGen

Digoxin: the Results of the DIG Study in the XXI Century

Izo Helber e Paulo J. F. Tucci

Escola Paulista de Medicina - Unifesp, São Paulo, SP - Brazil

Abstract

After the report that there was no statistical significance in the general mortality of the DIG study, the indication of digoxin in the treatment regimens for congestive heart failure (CHF) drastically decreased. *Post hoc* studies that reassessed the DIG study data, indicated that an aspect that was not considered in this multicenter study has a critical influence on the prognosis of patients: the serum levels of digoxin. Regarding those that received a placebo, the general mortality and hospitalization were decreased in patients with a digoxin level < 0.9 ng/ml. At the first study that assessed the influence of digitalis in an experimental model of CHF, we verified in

double-blind and placebo-controlled studies were published, which analyzed the clinical influence of the withdrawal of digoxin in patients that had been receiving the medication: PROVED³ and RADIANCE⁴. The patients followed at the PROVED trial received a diuretic associated to digoxin and the patients followed at the RADIANCE study received digoxin associated with diuretic and angiotensin-converting enzyme inhibitor. After the period of stabilization, the digitalis was substituted by placebo in one of the groups of each of these trials. In the two studies, weeks after the digoxin withdrawal, there was a decrease in exercise tolerance, decreased ejection fraction (EF), increased heart rate and decompensation in the

2. Arnold JMO, Liu P, Demers C, Dorian P, Giannetti M, Haddad H, et al. Canadian Cardiovascular Society consensus conference recommendations on heart failure 2006: diagnosis and management. *Can J Cardiol*. 2006; 22: 23-45.
3. Uretsky B, Young JB, Shahidi FE, Yellen LG, Harrison MC, Jolly MK. Randomized study assessing the effect of digoxin withdrawal in patients with mild to moderate chronic congestive heart failure: results of the Proved trial. *J Am Coll Cardiol*. 1993; 22: 955-62.
4. Packer M, Gheorghiade M, Young JB, Costantini DO, Adams KF Jr, Cody RJ, et al. Withdrawal of digoxin from patients with chronic heart failure treated with angiotensin-converting-enzyme inhibitors. *N Engl J Med*. 1993; 329: 1-7.
5. The Digitalis Investigation Group. The effect of digoxin on mortality and morbidity in patients with heart failure. *N Engl J Med*. 2007; 336: 525-33.
6. Eichhorn EJ, Gheorghiade M. Digoxin. *Prog Cardiovasc Dis*. 2002; 44: 251-66.
7. Eichhorn EJ, Lukas MA, Wu B, Shusterman N. Effect of concomitant digoxin and carvedilol therapy on mortality and morbidity in patients with chronic heart failure. *Am J Cardiol*. 2000; 86: 1032-5.
8. Pitt B, Zannad F, Remme WJ, Castaigne A, Perez A, Palensky J, et al. The effect of spironolactone on morbidity and mortality in patients with severe heart failure. Randomized Aldactone Evaluation Study Investigators. *N Engl J Med*. 1999; 341: 709-17.
9. Young JB, Gheorghiade M, Uretsky BF, Patterson JH, Adams KF Jr. Superiority of "triple" drug therapy in heart failure: Insights from the PROVED and RADIANCE trials. Prospective randomized study of ventricular function and efficacy of digoxin. randomized assessment of digoxin and inhibitors of angiotensin-converting enzyme. *J Am Coll Cardiol*. 1998; 32 (3): 686-92.
10. Gheorghiade M, Dirk J, Van Veldhuisen DJ, Colucci WS. Contemporary use of digoxin in the management of cardiovascular disorders. *Circulation* 2006; 114: 122-30.
11. Ahmed A. Digoxin and reduction in mortality and hospitalization in geriatric heart failure: importance of low doses and low serum concentration. *J Gerontol A Biol Sci Med Sci*. 2007; 62: 323-9.
15. Adams KF Jr, Patterson JH, Gattis WA, O'Connor CM, Lee CR, Schwartz TA, et al. Relationship of serum digoxin concentration to mortality and morbidity in women in the Digitalis Investigation Group Trial. A retrospective analysis. *J Am Coll Cardiol*. 2005; 46: 497-504.
16. Ahmed A, Gambassi G, Weaver MT, Young JB, Wehrmacher WH, Rich MW. Effects of discontinuation of digoxin versus continuation at low serum digoxin concentrations in chronic heart failure. *Am J Cardiol*. 2007; 100: 280-4.
17. Rathore SS, Curtis JP, Wang Y, Bristow MR, Krumholz HM. Association of serum digoxin concentration and outcomes in patients with heart failure. *JAMA*. 2003; 289: 871-8.
18. Ahmed A, Waagstein F, Pitt B, White M, Zannad F, Young JB, et al. Effectiveness of digoxin in reducing one-year mortality in chronic heart failure in the Digitalis Investigation Group trial. *Am J Cardiol*. 2009; 103: 82-7.
19. Ferguson DW, Berg WJ, Sanders JS, Roach PJ, Kempf JS, Kienzle MG. Sympathoinhibitory responses to digitalis glycosides in heart failure patients: direct evidence from sympathetic neural recordings. *Circulation*. 1989; 80: 65-77.
20. Gheorghiade M, Ferguson D. Digoxin: a neurohormonal modulator in heart failure? *Circulation*. 1991; 84: 2181-6.
21. Adams KF Jr, Gheorghiade M, Uretsky BF, Patterson JH, Schwartz TA, Young JB. Clinical benefits of low serum digoxin concentrations in heart failure. *J Am Coll Cardiol*. 2002; 39: 946-53.
22. Slatton ML, Irani WN, Hall SA. Does digoxin provide additional hemodynamic and autonomic benefit at higher doses in patients with mild to moderate heart failure and normal sinus rhythm? *J Am Coll Cardiol*. 1997; 29: 1206-13.
23. Pfeffer MA, Pfeffer JM, Steinberg C, Finn P. Survival after an experimental myocardial infarction: beneficial effects of long-term therapy with captopril. *Circulation*. 1985; 72: 406-12.

[Digoxin: the results of the X

Warning: Something's Not X

← → ↺

www.ncbi.nlm.nih.gov/pubmed/21180781

☆ 🔊 ☰

NCBI

Resources ▾ How To ▾

Sign in to NCBI

PubMed.gov

US National Library of Medicine
National Institutes of Health

PubMed ▾

Advanced

Search

Help

Display Settings: ▾ Abstract

Send to: ▾ free full text available at SciELO.org

Arq Bras Cardiol. 2010 Oct;95(4):e108-11.

[Digoxin: the results of the DIG study in the XXI century].

[Article in Multiple languages]

Helber I, Tucci PJ.

Author information

Abstract

After the report that there was no statistical significance in the general mortality of the DIG study, the indication of digoxin in the treatment regimens for congestive heart failure (CHF) drastically decreased. Post hoc studies that reassessed the DIG study data, indicated that an aspect that was not considered in this multicenter study has a critical influence on the prognosis of patients: the serum levels of digoxin. Regarding those that received a placebo, the general mortality and hospitalization were decreased in patients with a digoxin level < 0.9 ng/ml. At the first study that assessed the influence of digitalis in an experimental model of CHF, we verified in our lab that female rats with congestive syndrome secondary to myocardial infarction have a prolonged survival when undergoing treatment with digitoxin. The current information recommends that the merits of digoxin continue to be analyzed in order to adequately establish its importance in the treatment of CHF.

PMID: 21180781 [PubMed - indexed for MEDLINE] Free full text

Publication Types, MeSH Terms, Substances

LinkOut - more resources

PubMed Commons

0 comments

PubMed Commons home

Save items

☆ Add to Favorites ▾

Related citations in PubMed

Association of serum digoxin concentration and outcome [JAMA. 2003]

Review Digoxin remains useful in the management [Med Clin North Am. 2003]

[Effect of digoxin on atrial natriuretic peptide (ANI) [Pol Arch Med Wewn. 2001]

Review Digoxin use in congestive heart failure. Current status. [Drugs. 1998]

Review [The digitalis: should we forget it?]. [G Ital Cardiol (Rome). 2008]

See reviews

See all...

Related information

Related Citations

Compound (MeSH Keyword)

MedGen



Heart Failure

A heterogeneous condition in which the heart is unable to pump out sufficient blood to meet the metabolic need of the body. Heart failure can be caused by structural defects, functional abnormalities (VENTRICULAR DYSFUNCTION), or a sudden overload beyond its capacity. Chronic heart failure is more common than acute heart failure which results from sudden insult to cardiac function, such as MYOCARDIAL INFARCTION.

Year introduced: 2008 (1966)

PubMed search builder options

[Subheadings:](#)

- | | | |
|--|---|---|
| <input type="checkbox"/> analysis | <input type="checkbox"/> enzymology | <input type="checkbox"/> physiology |
| <input type="checkbox"/> anatomy and histology | <input type="checkbox"/> epidemiology | <input type="checkbox"/> physiopathology |
| <input type="checkbox"/> blood | <input type="checkbox"/> ethnology | <input type="checkbox"/> prevention and control |
| <input type="checkbox"/> cerebrospinal fluid | <input type="checkbox"/> etiology | <input type="checkbox"/> psychology |
| <input type="checkbox"/> chemically induced | <input type="checkbox"/> genetics | <input type="checkbox"/> radiography |
| <input type="checkbox"/> chemistry | <input type="checkbox"/> history | <input type="checkbox"/> radionuclide imaging |
| <input type="checkbox"/> classification | <input type="checkbox"/> immunology | <input type="checkbox"/> radiotherapy |
| <input type="checkbox"/> complications | <input type="checkbox"/> legislation and | <input type="checkbox"/> rehabilitation |
| <input type="checkbox"/> congenital | jurisprudence | <input type="checkbox"/> statistics and numerical |
| <input type="checkbox"/> cytology | <input type="checkbox"/> metabolism | data |
| <input type="checkbox"/> diagnosis | <input type="checkbox"/> microbiology | <input type="checkbox"/> surgery |
| <input type="checkbox"/> diet therapy | <input type="checkbox"/> mortality | <input type="checkbox"/> therapy |
| <input type="checkbox"/> drug therapy | <input type="checkbox"/> nursing | <input type="checkbox"/> ultrasonography |
| <input type="checkbox"/> economics | <input type="checkbox"/> organization and | <input type="checkbox"/> urine |
| | administration | <input type="checkbox"/> veterinary |

Filters: [Manage Filters](#)

[Show additional filters](#)

Related citations

6083 free full-text articles in PubMed Central

["congestive heart failure"](#) X

[www.ncbi.nlm.nih.gov/pubmed/?term="congestive+heart+failure"+AND+digoxin+AND+hospital*](#)

[NCBI](#)
[Resources](#)
[How To](#)
[Sign in to NCBI](#)

[PubMed.gov](#)
 US National Library of Medicine
 National Institutes of Health

PubMed
 "congestive heart failure" AND digoxin AND hospital*
 Search

[RSS](#)
[Save search](#)
[Advanced](#)
[Help](#)

[Show additional filters](#)

Display Settings:
☒ Summary, 20 per page, Sorted by Recently Added

Send to:
☒
Filters:
[Manage Filters](#)

Article types

Clinical Trial

Review

More ...

Text availability

Abstract available

Free full text available

Full text available

Publication dates

5 years

10 years

Custom range...

Species

Humans

Other Animals

[Clear all](#)

[Show additional filters](#)

Wildcard search for 'hospital*' used only the first 600 variations. Lengthen the root word to search for all endings.

Results: 1 to 20 of 323

<< First
 < Prev
 Page 1 of 17
 Next >
 Last >>

- ☐
[Evaluation of the concurrent use of potassium-influencing drugs as risk factors for the development of hyperkalemia.](#)

1. Kuijvenhoven MA, Haak EA, Gombert-Handoko KB, Crul M.
 Int J Clin Pharm. 2013 Dec;35(6):1099-104. doi: 10.1007/s11096-013-9830-8. Epub 2013 Aug 22.
 PMID: 23974985 [PubMed - in process]
[Related citations](#)

- ☐
[Influences of simvastatin on vascular endothelial function of patients with coronary heart disease complicated with congestive heart failure.](#)

2. Wang L, Shi J, Zhang Y.
 Eur Rev Med Pharmacol Sci. 2013 Jun;17(12):1590-3.
 PMID: 23832723 [PubMed - indexed for MEDLINE] **Free Article**
[Related citations](#)

- ☐
[Digoxin Downregulates NDRG1 and VEGF through the Inhibition of HIF-1α under Hypoxic Conditions in Human Lung Adenocarcinoma A549 Cells.](#)

3. Wei D, Peng JJ, Gao H, Li H, Li D, Tan Y, Zhang T.
 Int J Mol Sci. 2013 Apr 2;14(4):7273-85. doi: 10.3390/ijms14047273.
 PMID: 23549264 [PubMed] **Free PMC Article**
[Related citations](#)

- ☐
[Rhythm versus rate control therapy and subsequent stroke or transient ischemic](#)

OTHER WEBSITES



**NATIONAL GUIDELINE CLEARINGHOUSE
THE COCHRANE COLLABORATION
NEW ENGLAND JOURNAL OF MEDICINE
SPECIALTY JOURNALS**

Once you find an article,
What will you do with it?



Do the Critical Appraisal!



USE

- The Evidence-Based Family and Community Practice (EBFCP) and Quality Improvement in Health Care Manual (FMRG, 2003)

Other source to Understand:

- Painless EBM by Dr. Dans
- World Wide Web
- BUT the guide questions to be answered must come from the EBFCP Manual

FORMAT OF JOURNAL REPORT



- Case Scenario
- Research Question
- Search
- Title
- Source
- Authors
- Appraisal

EVIDENCE-BASED FAMILY AND COMMUNITY PRACTICE



- I. Is it relevant?
- II. Is it valid?
- III. What are the results?
- IV. Is it applicable to my patient?

Critical Appraisal of An Article on Diagnostics



Decision analysis



- appraisal of an article on diagnostics start with the decision analysis line

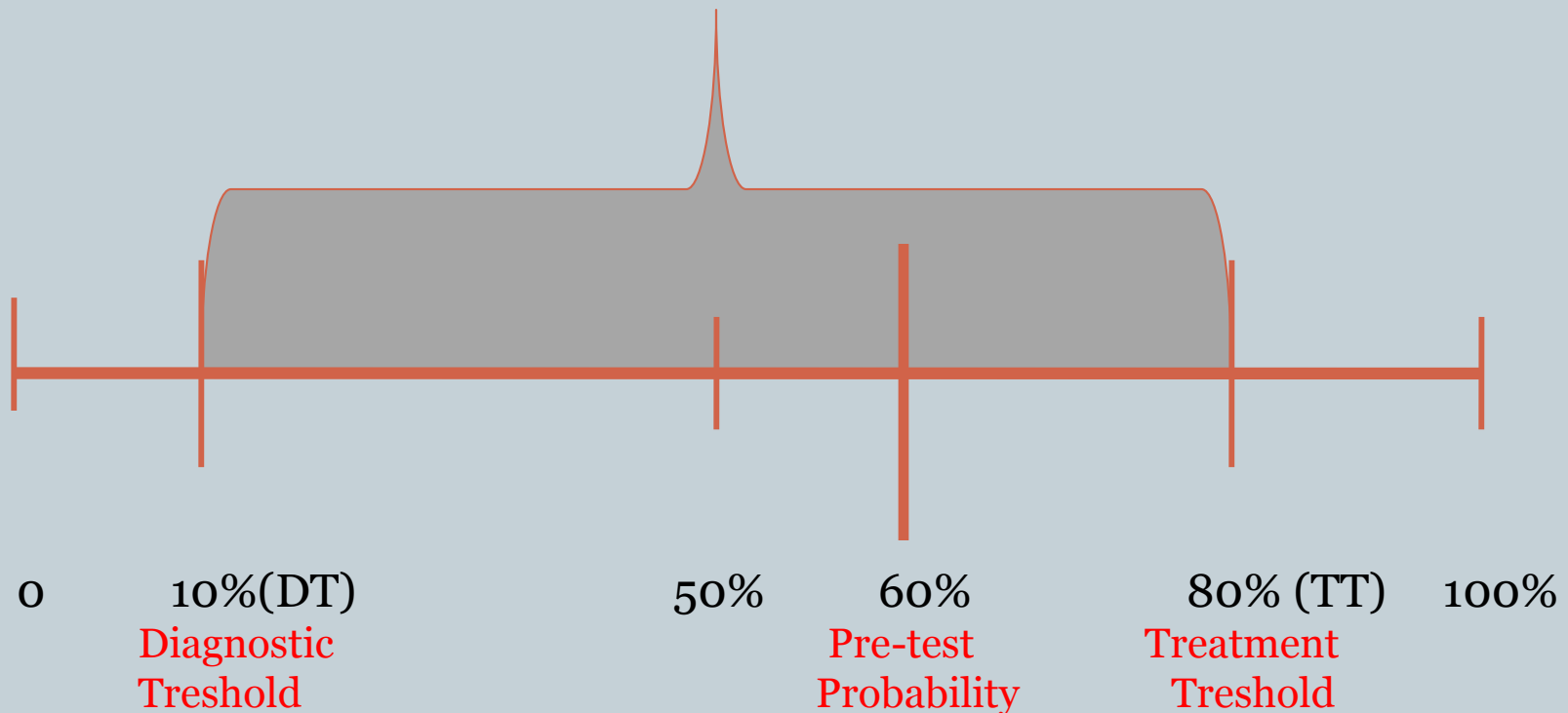
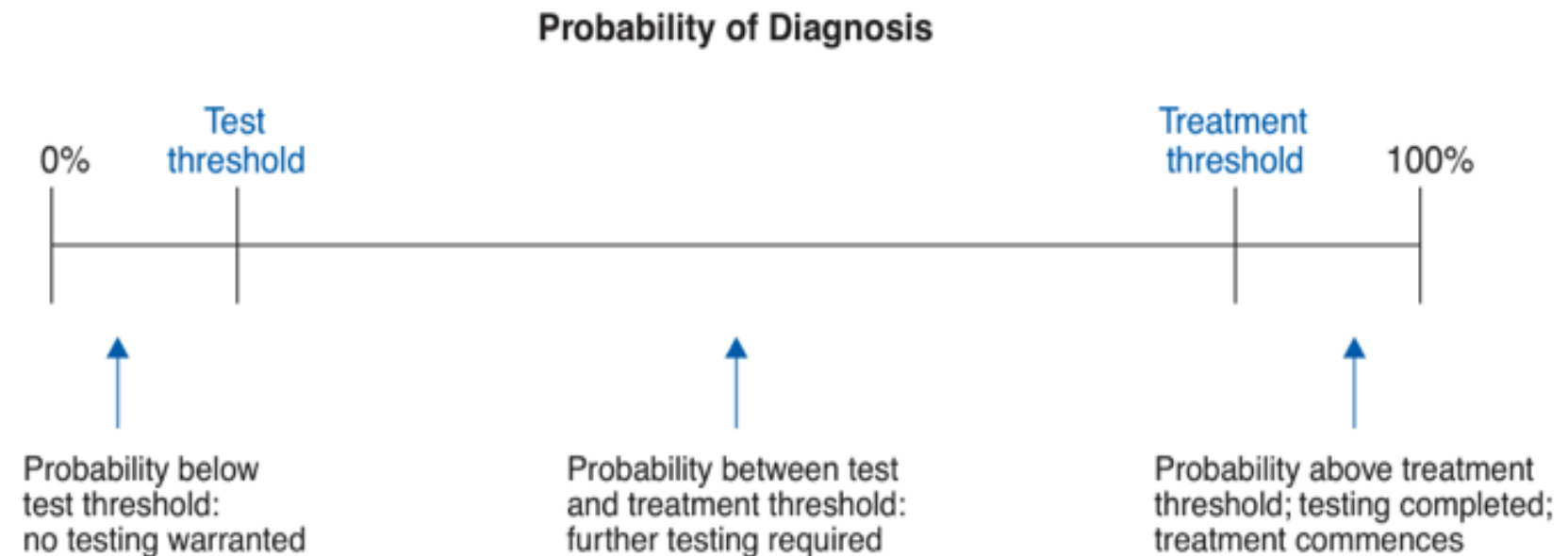


FIGURE 14-2

Test and Treatment Thresholds in the Diagnostic Process



Decision Analysis Line



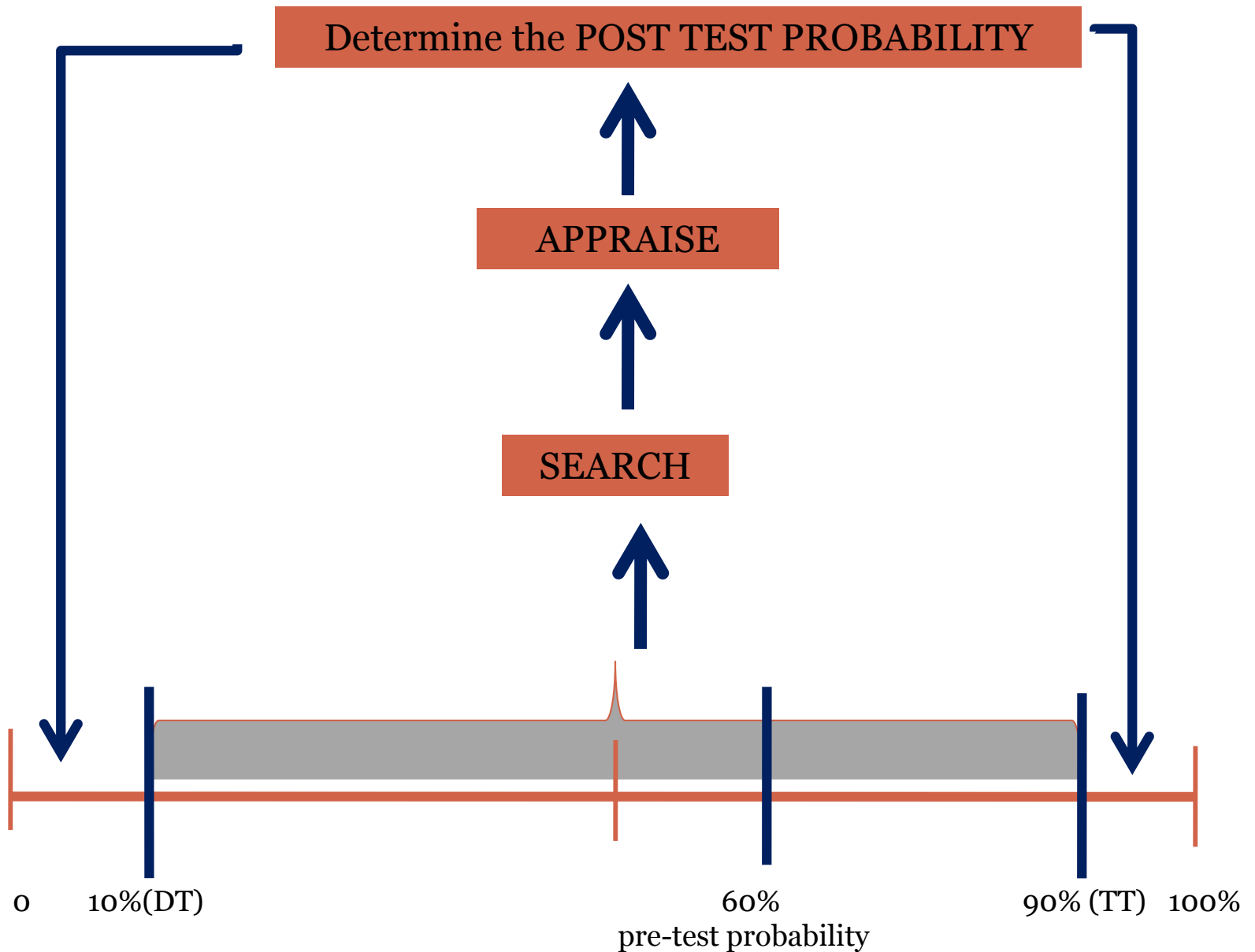
- Diagnostic Threshold (DT): arbitrary point at which you rule out the disease below this point
 - After this point, you will request for a diagnostic test
- Therapeutic Threshold: arbitrary point at which you decide to treat beyond this point
 - Below this point, you will request for a diagnostic test
- Pre-Test Probability: probability that the patient has the disease
- Post-Test probability: probability that the patient has the disease after doing the diagnostic exam

Case Scenario



- 24 year old male
- Presenting with 1 month history of RUQ pain
- Started out as epigastric pain then localizing in the RUQ
- Usually associated with intake of fatty food
- No other associated symptom
- PE: (-) Murphy's

- What is our pre test probability that this patient has Acute Cholelithiasis?
- Will we request for an HBT ULTRASOUND?



Is it relevant?



- Is the objective of the study similar to your clinical dilemma?

	CLINICAL DILEMMA	STUDY
Population		
Intervention and comparative intervention	?	?
Methodology	?	?
Outcome		

Is it valid?



PRIMARY VALIDITY GUIDES

1. Was there an independent comparison with a reference standard?
2. Did the patient sample include an appropriate spectrum of patients to whom the test will be done?

SECONDARY VALIDITY GUIDES

1. Was the reference standard done regardless of the result of the of the diagnostic test being evaluated?
2. Were the methods for performing the test described in sufficient detail to permit replication?

PRIMARY VALIDITY GUIDES



1. Was there an independent comparison with a reference standard?

Reference standard.

Accuracy.

Precision.

* The reference standard should not be part of the diagnostic procedure in question.

PRIMARY VALIDITY GUIDES



2. Did the patient sample include an appropriate spectrum of patients to whom the test will be done?

- Inclusion and Exclusion Criteria.
- Baseline Characteristics (table 1).
- Representativeness includes subjects with the whole spectrum of the disease
- The accuracy of a diagnostic test among patients with low risk for the disease is different from patients with high risk for the disease.

SECONDARY VALIDITY GUIDES



1. Was the reference standard done regardless of the result of the of the diagnostic test being evaluated?

SECONDARY VALIDITY GUIDES



2. Were the methods for performing the test described in sufficient detail to permit replication?

2. Were the methods for performing the test described in sufficient detail to permit replication?

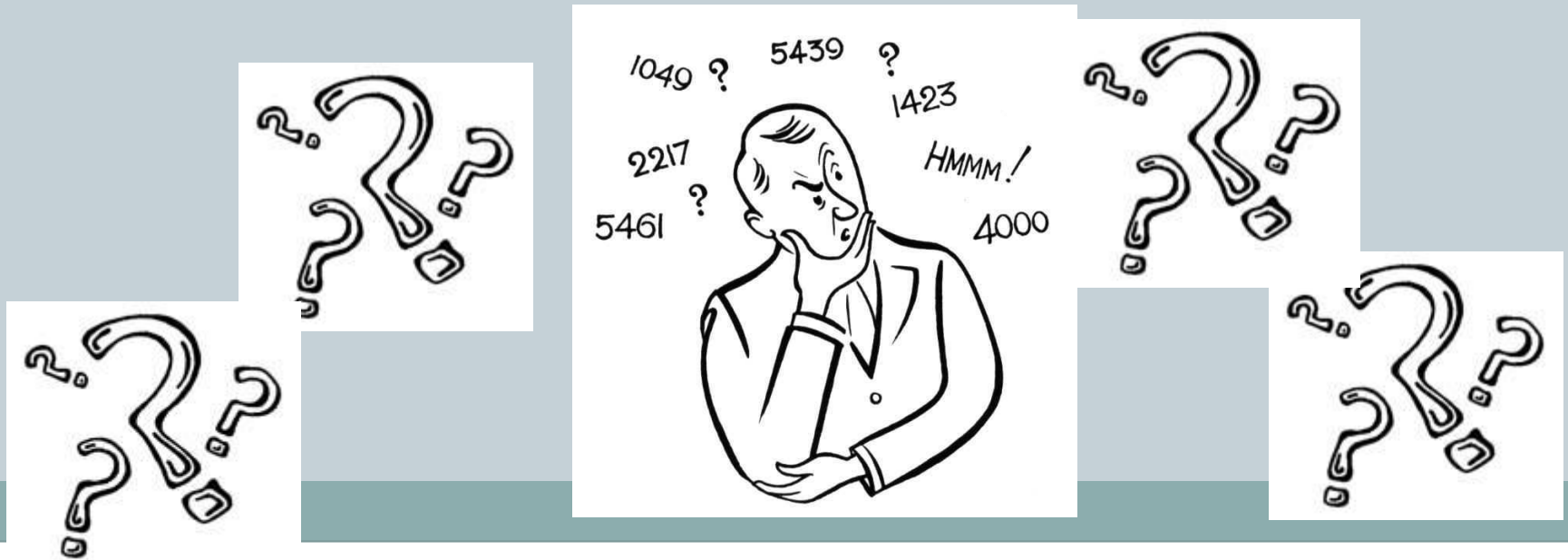


- Look at the results and methodology section.
- Clear procedure including the preparation of subjects.
 - Diets, drugs to avoid.
 - Precautions.
 - Step by step descriptions.
- Be able to duplicate the exam in your setting and still get the same results.

II. Overall is the study valid?



- If you want to be strict about it, you should answer yes in all 4 questions.
- A simple rule might be to answer yes to at least, one primary guide and two secondary guides.



III. What are the results?



What are the likelihood ratios for the different test results?

Likelihood ratios indicates by how much a given test result increases the pre-test probability of the disease.

LR of 1 means that the pre-test probability is same after doing the test

LR >1 increases the probability that the disease is present

Sensitivity & Specificity



Disease

Test

	Present	Absent
Positive	TP	FP
Negative	FN	TN

SENSITIVITY: probability/likelihood that the diseased patient will test positive: $\text{TP} / \text{TP} + \text{FN}$

SPECIFICITY: probability/likelihood that those without the disease will test negative: $\text{TN} / \text{TN} + \text{FP}$

Remember? **SpPIN & SnNOUT**

LIKELIHOOD RATIO

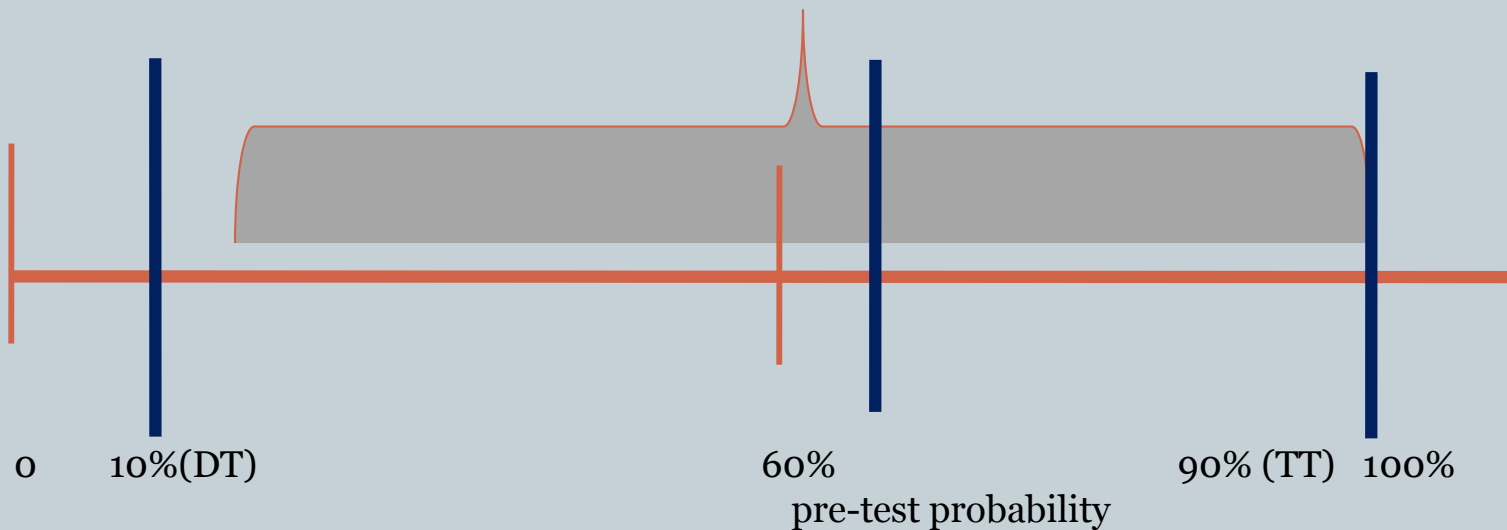


- Likelihood ratio of a positive test
 - Probability that the test is + among diseased person
Probability that the test is + among non-diseased person
 - $LR (+) = \text{Sensitivity} / 1 - \text{Specificity}$
- Likelihood ratio of a negative test
 - Probability that the test is - among diseased person
Probability that the test is - among non-diseased person
 - $LR (-) = 1 - \text{Sensitivity} / \text{Specificity}$

What do Likelihood ratios do?

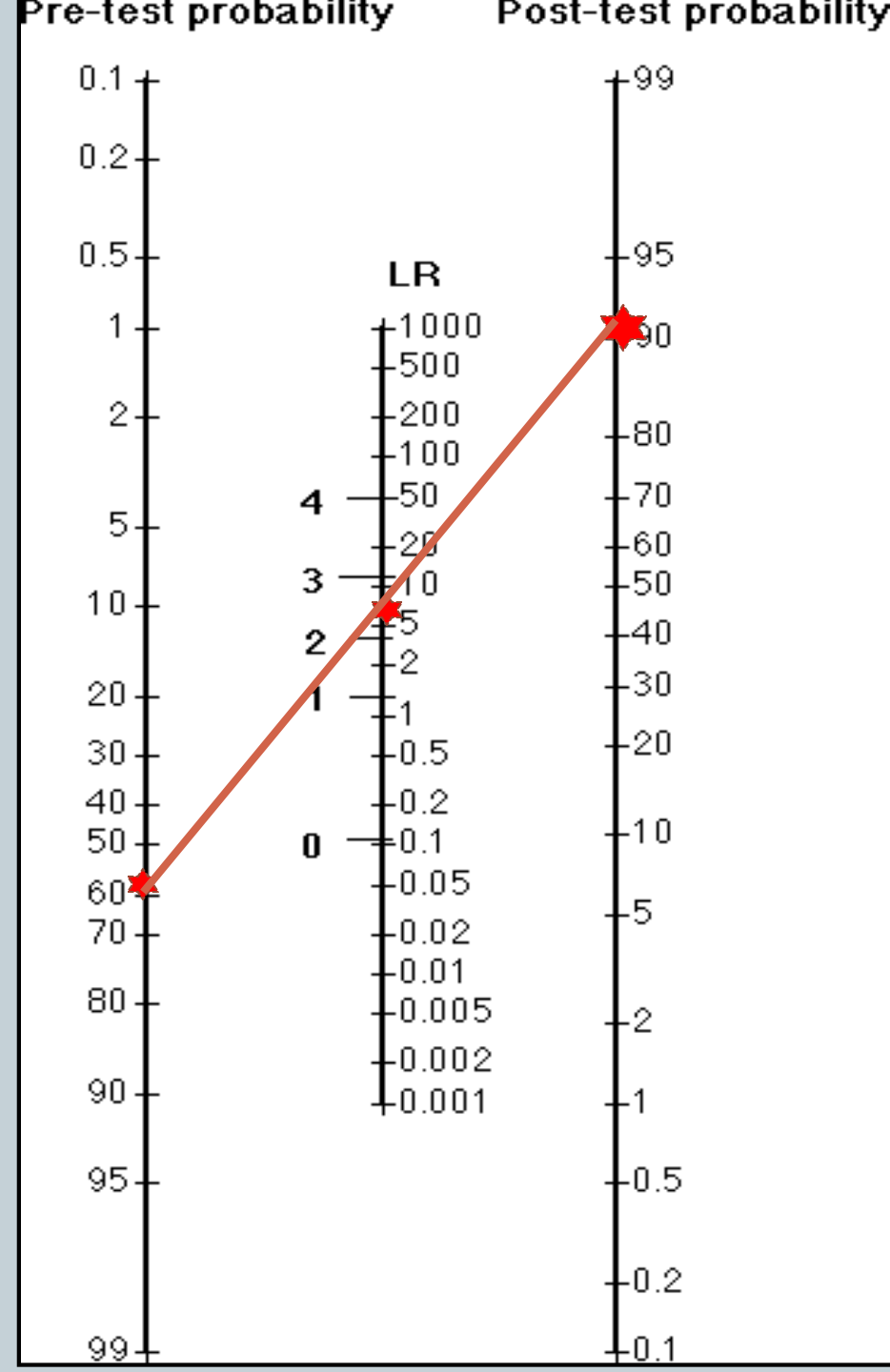


- It can change the probability of the disease after a diagnostic test is done.
- Remember our Case Scenario



Will the likelihood ratios shift the pre-test probability

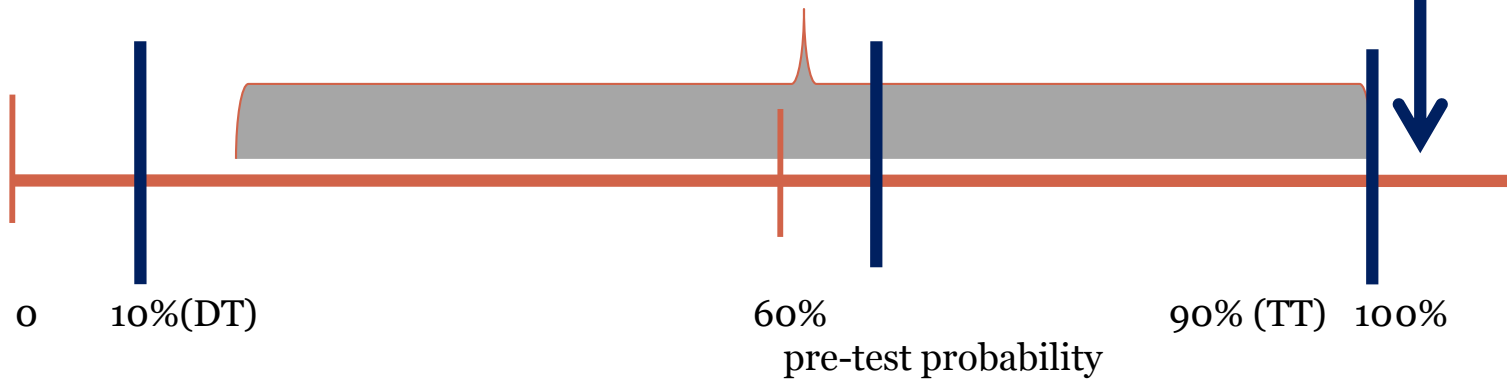
- Use a NOMOGRAM
- Given a pre test probability of 60%
- The HBT UTZ yielded a LR of 2.5
- PLOT
- Post test Probability is 93%
- What does this mean?



Determine the POST TEST PROBABILITY

APPRAISE

SEARCH



IV. Can the results help me in caring for my patients?



1. Will the reproducibility of the test result and its interpretation be satisfactory in my setting?
 - Is the interpretation simple enough?
 - Is the basis of the interpretation clear and specific?
2. Are the results applicable to my patient?
 - setting is somewhat similar
 - inclusion criteria include characteristics of your patient
 - clinical judgment is required
3. Will the results change my management?

Critical appraisal of an article on therapeutics



Is it relevant?



- Is the objective of the study similar to your clinical dilemma?

	CLINICAL DILEMMA	STUDY
Population		
Intervention and comparative intervention	?	?
Methodology	?	?
Outcome		

I. Is it valid?



Primary Validity Guides

1. Was the assignment of patients to treatment randomized?
2. Were all the patient who entered the trial properly accounted for and attributed at its conclusion?

Secondary Validity Guides

1. Were patient, their clinicians and study personnel “blinded”?
2. Were the groups similar at the start of the trial?
3. Aside from the experimental intervention, were the groups treated equally?

1. Was the assignment of patients to treatment randomized?



- Look at the abstract or methodology.
- Define randomization.
- Random sampling?
- Randomization assures that both known and unknown determinants of outcome are even distributed between the treatment and control
- Avoids selection bias
- More severe disease = worse results
- less severe disease = better results

2. Were all the patient who entered the trial properly accounted for and attributed at its conclusion?



b. Were the patients analyzed in the group to which they were randomized?

- patients, including drop outs or withdrawn, were analyzed to their original grouping at the start of the study
- This preserves the value of randomization
- No crossing over

2. Were all the patient who entered the trial properly accounted for and attributed at its conclusion?



a. Was follow up complete?

- Methodology and Results Section
- Look at the number of patients enrolled at the outset and compare this with the number of patient reported at the results table.
- What is the drop out rate?
- 20 % or more is considered substantial
- If less than 20% dropped out, check if and *intention to treat analysis* was done
- What is intention to treat analysis?

I. Is it valid?



Primary Validity Guides

1. Was the assignment of patients to treatment randomized?
2. Were all the patient who entered the trial properly accounted for and attributed at its conclusion?

Secondary Validity Guides

1. Were patient, their clinicians and study personnel “blinded”?
2. Were the groups similar at the start of the trial?
3. Aside from the experimental intervention, were the groups treated equally?

1. Were patients, their clinicians, and study personnel "blind" to treatment?



- Look at the abstract or methodology.
- Define blinding.
- What is its importance?
 - SINGLE blinding?
 - DOUBLE blinding?
 - TRIPLE blinding?

2. Were the groups similar at the start of the trial?



- Baseline characteristics usually labeled

Table no. 1

- The greater the similarity between known prognostic factors for the control and experimental group, the more likely that the results can be attributed to the intervention, rather than due to the differences in these factors

3. Aside from the experimental intervention, were the groups treated equally?

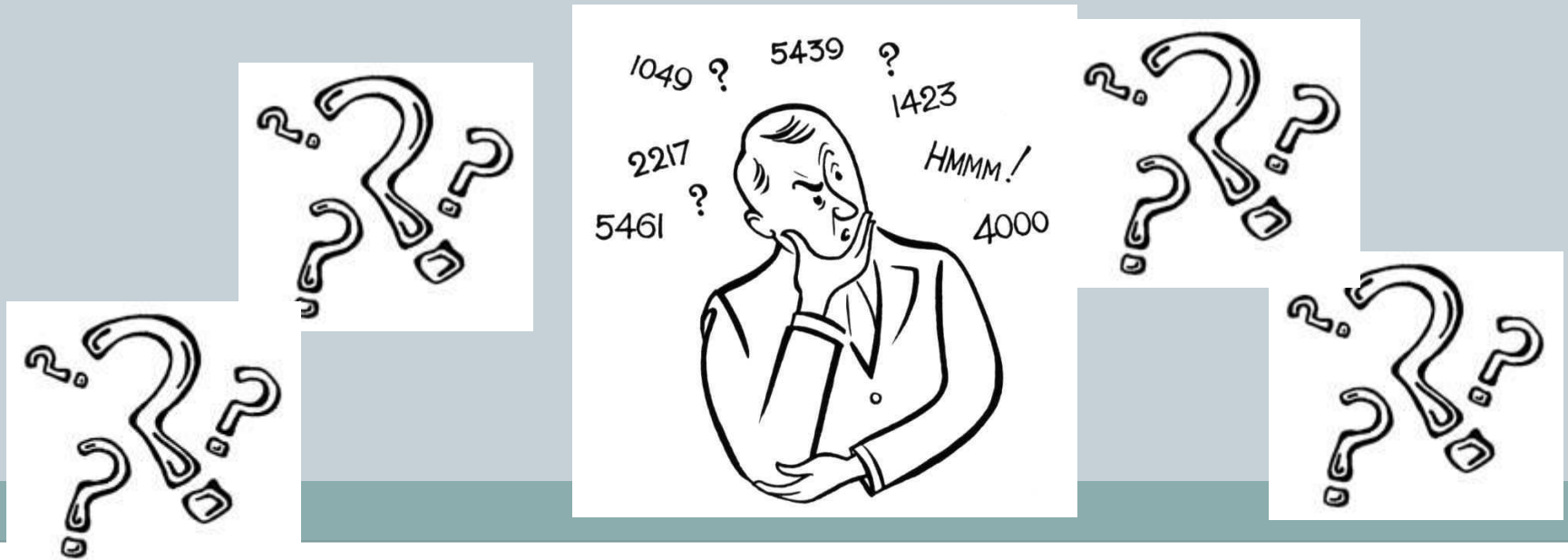


- Look for interventions other than the treatment under study
- “co-interventions” distort the results
- Where outcomes measurements clearly described and determined in the same way between interventions?
- **DEFINE**
 - Clinical Outcomes
 - Surrogate Outcomes

II. Overall is the study?



- If you want to be strict about it, you should answer yes in all 5 questions.
- A simple rule might be to answer yes to at least, one primary guide and two secondary guides.



III. WHAT ARE THE RESULTS?



- a. How large was the treatment effect?

RISK IN TREATMENT (R_t)

$$\frac{\text{No. of patients who did not get well in the treatment group}}{\text{Total no. of patients in the treatment group}}$$

RISK IN CONTROL (R_c)

$$\frac{\text{No. of patients who did not get well in the control group}}{\text{Total no. of patients in the control group}}$$

a. What are the results?



- Absolute Risk Reduction (**ARR**) = $R_c - R_t$
- Relative Risk (**RR**) = R_t / R_c

RR of 1 : No difference between Treatment and Control

RR of >1: Treatment is more harmful

RR of <1: Treatment is more effective

- Relative Risk Reduction (**RRR**) = $1 - RR$

B. How precise was the treatment effect?



- What is the confidence interval?
- What is the p-value?

IV. Are the results applicable to my patients?



- Are the medical, social and economic resources needed to administer the treatment available in your setting?
- Consider the acceptability by your patient, his family and community.
- Are the likely treatment benefits worth the potential harm and costs?

Number needed to treat – $1/ARR$
Cost effectiveness Formula: $NNT \times \text{price/unit} \times \text{dose} \times \text{duration}$

THANK YOU!

**The doctor of the future
will give no medicine, but
will interest her or his
patients in the care of the
human frame, in a proper
diet, and in the cause and
prevention of disease.**

Thomas A. Edison
US inventor (1847 - 1931)