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What 20 Years of Research on Diagnosis Have Taught Me (that others should know)

Dr Olga Kostopoulou

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#SCALearning25

IMPERIAL

**What 20 years of research on diagnosis
have taught me (that others should know)**

Olga Kostopoulou, PhD
Reader in Medical Decision Making

Diagnostic error

Overlooked in patient safety?

- “*Diagnosis apparently gets overlooked in most efforts to ensure quality and safety*” (Leape et al., JAMA 2002)
- IoM report 1998 “To Err Is Human”: the term *medication error* was mentioned 70 times, while *diagnostic error* appeared only twice.
- IoM report 2015 “Improving Diagnosis in Health Care”: “*Most people will experience at least one diagnostic error in their lifetime*”.

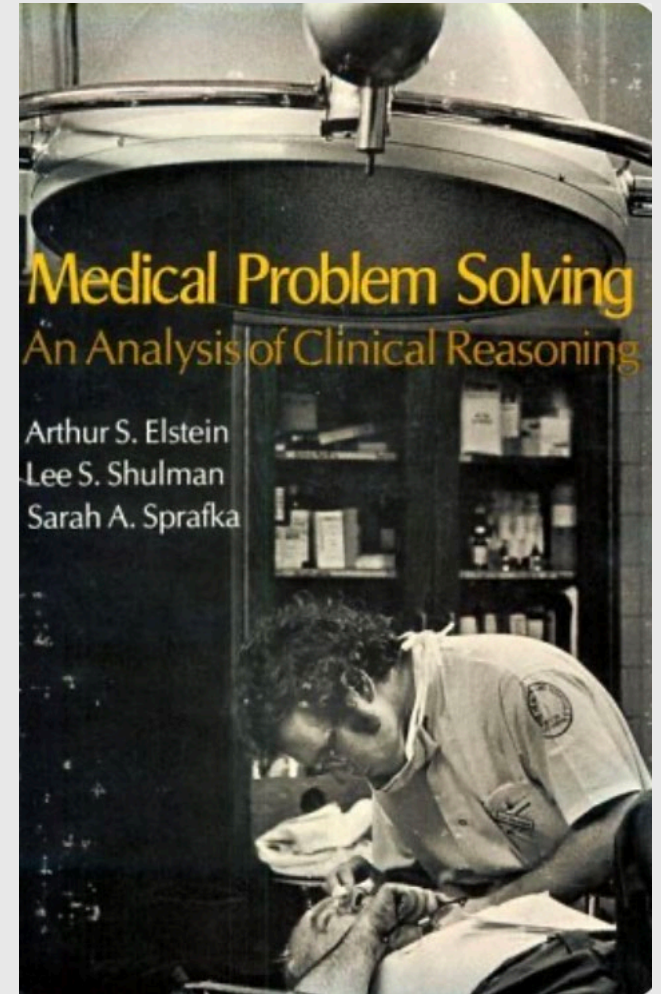
Diagnostic error

A Cognition-System interplay

- Cognitive causes are common (Gandhi et al., 2006; Lyratzopoulos, et al., 2015)
- Cognition is relevant to essentially every aspect of diagnosis and medical practice generally (Croskerry & Graber, 2025)
- E.g.,
 - Failing to consider a broad enough differential
 - Focusing on one diagnosis too early
 - Not asking enough questions
 - Dismissing things that don't 'fit'
- *Not specific to medicine*

Ample evidence that

- People formulate hypotheses quickly and with little information (Baron, 2000)
- Automatically, even if they are instructed not to do so. *A psychological necessity.*
- Clinicians generate diagnostic hypotheses within seconds (Elstein et al. 1978)
- Diagnostic hypotheses guide information search and interpretation



Diagnosis as a cognitive process

- Where do diagnostic hypotheses come from?

Memory.

- How do they get activated?

From observations (e.g., symptoms, signs, risk factors, FHx, GP's referral letter, etc.)

- Which hypotheses will be activated first?

Those that are more strongly linked with the observations (common diagnoses, serious diagnoses)

- People will generate only a small subset of hypotheses - no more than 4.

First impressions matter

And clinicians know it

- Physicians acknowledge their importance in interviews (e.g., Balla et al. 2012; Bösner et al. 2014)
- the first impression ‘...is **crucial** regarding the decision on what I will do with this patient and on what further diagnostic procedures I will initiate.’ (Bösner et al. 2014)

Initial hypotheses are compelling

- Rested working memory
- More attention, better encoding
- Disproportionate weight ('primacy effect')
- Crowd out competitors
- The lens through which incoming information is evaluated

First impressions matter

How much?



Kostopoulou et al. (2017) *Medical Decision Making*, **37**(1), 9-16.

ORIGINAL ARTICLE

The Role of Physicians' First Impressions in the Diagnosis of Possible Cancers without Alarm Symptoms

*Olga Kostopoulou, PhD, Miroslav Sirota, PhD, Thomas Round, MBBS,
Shyamalee Samaranayaka, MD, Brendan C. Delaney, MD*

- Are GPs' initial diagnostic impressions associated with subsequent diagnosis and referral decisions?
- Study aimed to elicit and measure this association

Design

- 90 UK GPs
- 6 detailed patient scenarios (3 possible cancers)
 - Repeated consultations
- Interactive via phone communication with researcher

Henry Strauss

Patient information

- AGE: 61 years old
- ETHNICITY: Caucasian
- HEIGHT: 175 cm (5ft 7.7")
- WEIGHT: 72 kg (11.3 stones) (BMI 23.5, measured 6 months ago)
- SMOKING STATUS: Never smoked
- ALCOHOL: Occasional
- LAST BLOOD PRESSURE READING: 138/82, taken 6 months ago
- PAST MEDICAL HISTORY: Irritable Bowel Syndrome (2000) - coeliac screening negative. Hypertension (2006).
- MEDICATIONS: Lisinopril 5 mg od
- OCCUPATION: Administrative officer in the local Council, now retired
- LAST CONSULTATION: 6 months ago for routine review of hypertension
- APPEARANCE: Nothing of note.

Presenting complaint

Doctor, I am having problems with my bowels again. I've been having hard stools for about a month now. I go to the loo only once in 4-5 days.

Confirm you have read the Presenting Complaint

Henry Strauss

Patient information

- AGE: 61 years old
- ETHNICITY: Caucasian
- HEIGHT: 175 cm (5ft 7.7")
- WEIGHT: 72 kg (11.3 stones) (BMI 23.5, measured 6 months ago)
- SMOKING STATUS: Never smoked
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- APPEARANCE: Nothing of note.

Hb 13.0 g/dL (13.5-18)*

MCV 76 fL (76-100)

WCC 8.1×10^9 (4.0-11.0)

Neutrophils 6.2×10^9 (2.0-7.5)

Lymphocytes 2.0×10^9 (1.3-3.5)

Monocytes 0.2×10^9 (0.2-0.8)

Eosinophils 0.3×10^9 (0-0.4)

Platelets 295×10^9 (150-450)

Two weeks later, Henry Strauss comes back to see you.

Patient information

- AGE: 61 years old
- ETHNICITY: Caucasian
- HEIGHT: 175 cm (5ft 7.7")
- WEIGHT: 72 kg (11.3 stones) (BMI 23.5 measured 6.5 months ago)
- SMOKING STATUS: Never smoked
- ALCOHOL: Occasional
- LAST BP: 132/78, taken 2 weeks ago
- PAST MEDICAL HISTORY: Irritable Bowel Syndrome (2000) - coeliac screening negative. Hypertension (2006).
- MEDICATION: Lisinopril 5 mg od
- OCCUPATION: Administrative officer in the local Council, now retired
- LAST CONSULTATION: 2 weeks ago with constipation for one month.
- APPEARANCE: Nothing of note.

Presenting complaint

Good morning doctor. Do you remember that I came to see you two weeks ago with constipation? I did everything you told me but it hasn't gone away. I'm not sure what else to do.

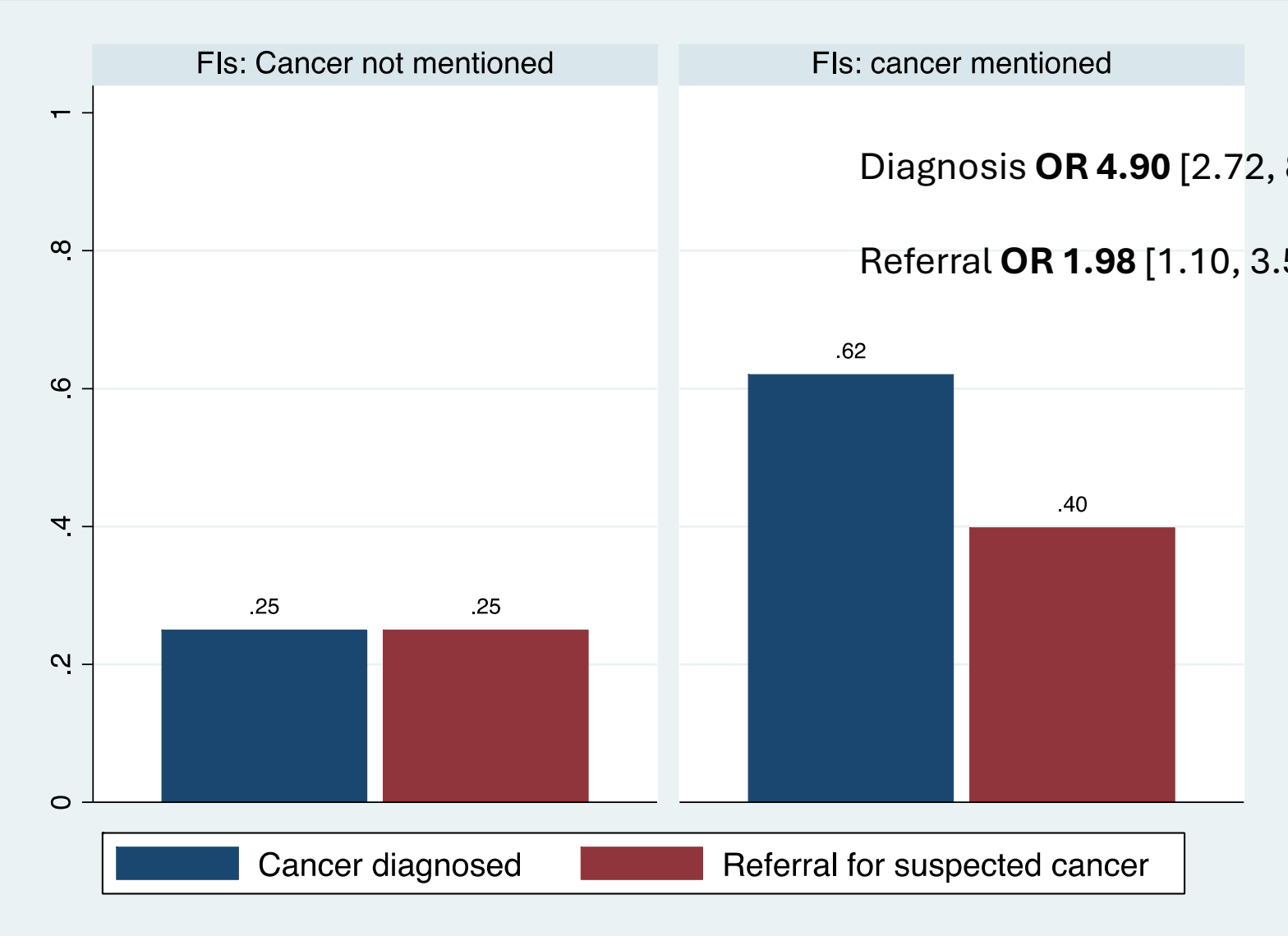
Confirm you have read the Presenting Complaint

Design

- Think aloud
- Coded and analysis of initial utterances
 - Cancer mentioned? Yes (1), No (0)

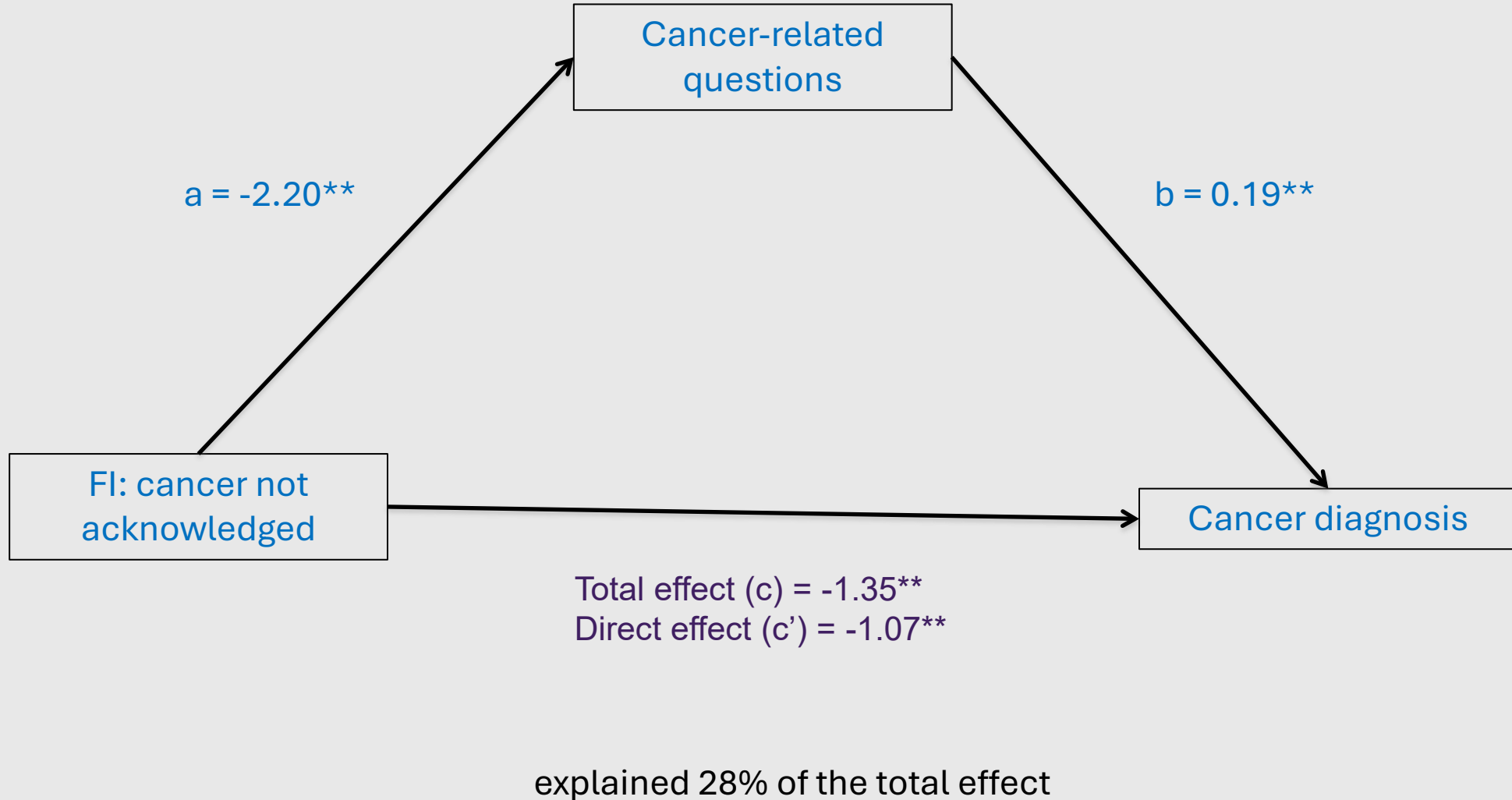
Early, explicit acknowledgement of cancer possibility

Significantly increased cancer diagnosis and referral odds



First impressions drive information gathering

A mediation model



First impressions drive interpretation

Predecisional Information Distortion (PID)

- We tend to evaluate information in a biased way in order to:
 1. Consolidate a decision already made (post-decision)
 2. Strengthen an emerging decision (pre-decision)
- Incoming information is evaluated as supportive of an emerging judgement, preference or currently leading hypothesis (Russo et al. 1998; DeKay, 2015)
- Innate drive to attain and maintain ‘cognitive consistency’ (Russo et al. 2008; Simon et al. 2004).

PID in clinical diagnosis

Medical Decision Making (2012) **32**(6), 831-839

Information Distortion in Physicians' Diagnostic Judgments

*Olga Kostopoulou, PhD, J. Edward Russo, PhD, Greg Keenan, MSc,
Brendan C. Delaney, MD, Abdel Douiri, PhD*

Judgment and Decision Making, Vol. 9, No. 6, November 2014, pp. 572–585

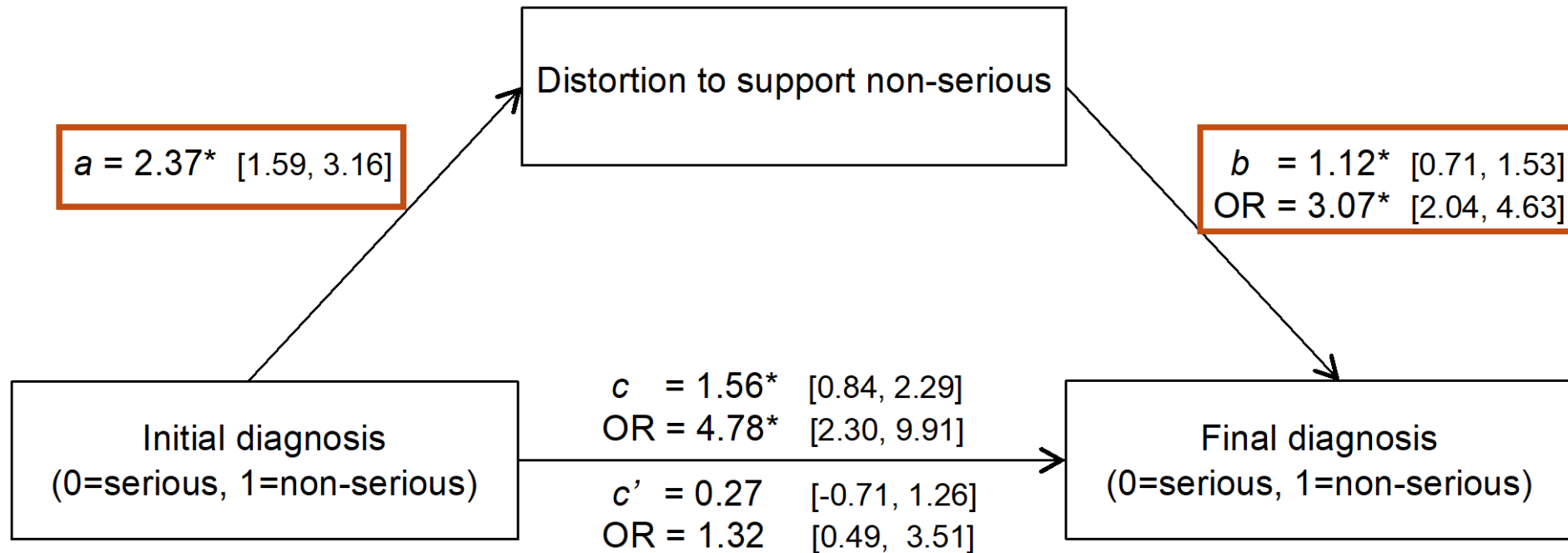
**Predecisional information distortion in physicians' diagnostic
judgments: Strengthening a leading hypothesis or weakening its
competitor?**

Martine Nurek*

Olga Kostopoulou[†]

York Hagmayer[‡]

PID explained 91% of the total effect c



* $P < 0.001$

Confidence can increase bias

- The more certain we are about an initial hypothesis, the less likely we are to test it sufficiently and rigorously

ORIGINAL ARTICLE

Open Access

Influences of early diagnostic suggestions on clinical reasoning



Ploutarchos Kourtidis, Martine Nurek, Brendan Delaney and Olga Kostopoulou* 

“Initial certainty was the main driver of behaviour: high certainty led to significantly fewer information requests, more biased information evaluations, and fewer changes in diagnosis when encountering new information that either suggested additional possibilities or did not entirely fit with one’s leading hypothesis. The negative relationship between initial certainty and change in diagnosis was **consistent across the three experiments**” (N s=194, 248, 190)

Individual differences in thinking style

Actively Open-minded Thinking (AOT)

- Are you **open** to alternative explanations that oppose an initial judgement?
- Do you **actively** search for evidence to disconfirm pre-established beliefs (Baron, 2006, 2019).
- Example AOT questions:
 - People should revise their conclusions in response to relevant new information.
 - Changing your mind is a sign of weakness.
 - People should search actively for reasons why they might be wrong.

AOT and diagnosis

- Active open-mindedness was associated with more requests for information, more investigations, and more differential diagnoses (Kourtidis et al. 2022).



Debiasing thinking

Good thinking strategies

To increase AOT

- Facilitate consideration of alternatives
 - Direct attention to evidence that would otherwise not be considered.
1. Consider the opposite (What if...?)
 2. Self-generate alternatives
 3. Counteract unjustified confidence (Why might I be wrong? Consider reasons why my initial judgement might be wrong)
 4. Designate a devil's advocate (human or AI)
 5. Adopt another stakeholder's perspective (patient, carer, medical director)

Do these work in medicine?

- Scarce evidence, mostly negative (no impact)
 - Medical students or junior doctors
 - Stand-alone, one-off interventions
 - Written vignettes
- Can they be deployed spontaneously?

Jess's Rule

'Three strikes and we RETHINK'

R

Review the case thoroughly after three unresolved or escalating presentations.

E

Elevate the concern for a 'fresh eyes' approach and clinical reassessment.

T

Think again—especially if the original diagnosis was unsubstantiated.

H

Hear the patient fully and consider what's changed or been missed.

I

Invite a second opinion or peer discussion within Primary Care.

N

Navigate referral to secondary care where appropriate.

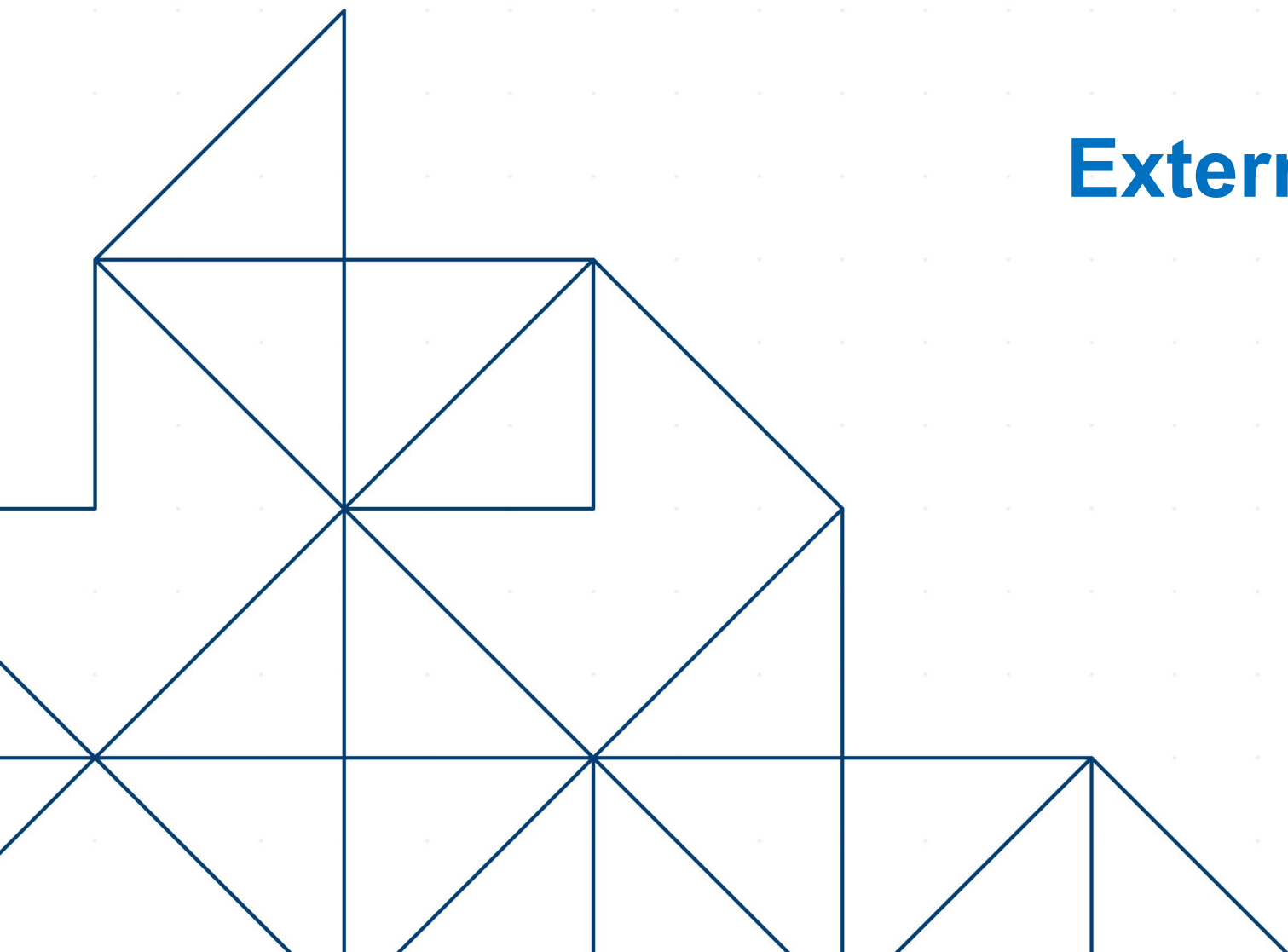
K

Keep continuity with the same GP and ensure in-person assessment if previous care was remote.



Royal College of
General Practitioners





External aids

Diagnostic Decision support systems

Differential diagnosis generators

- Potentially helpful – mixed evidence
- Small but significant improvement in accuracy (before-after studies)
- Poor methodological quality of the included studies & high between-studies heterogeneity

- Extensive lists of low relevance

- Inconclusive evidence re: number of investigations, cost-effectiveness



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RESEARCH ARTICLE

The Effectiveness of Electronic Differential Diagnoses (DDX) Generators: A Systematic Review and Meta-Analysis

Nicholas Riches , Maria Panagioti, Rahul Alam, Sudeh Cheraghi-Sohi, Stephen Campbell, Aneez Esmail, Peter Bower

Published: March 8, 2016 • <https://doi.org/10.1371/journal.pone.0148991>

Potential biasing effect

- “when the most likely diagnosis was included in the list of suggestions (vs. not included), physicians who gave that diagnosis initially, tended to request less information, evaluate it as more supportive of their diagnosis, become more certain about it, and change it less frequently when encountering new but ambiguous information; in other words, they seemed to **validate rather than question** their initial hypothesis.”

Kourtidis et al.
Cognitive Research: Principles and Implications (2022) 7:103
<https://doi.org/10.1186/s41235-022-00453-y>

Cognitive Research: Principles
and Implications

ORIGINAL ARTICLE

Open Access

Influences of early diagnostic suggestions on clinical reasoning



Ploutarchos Kourtidis, Martine Nurek, Brendan Delaney and Olga Kostopoulou* 

Potential biasing effect

- “The look at the ddx list seems to serve only one purpose: the verification of the diagnostic assumption” (p. 98).



ELSEVIER

International Journal of Medical Informatics 53 (1999) 91–100

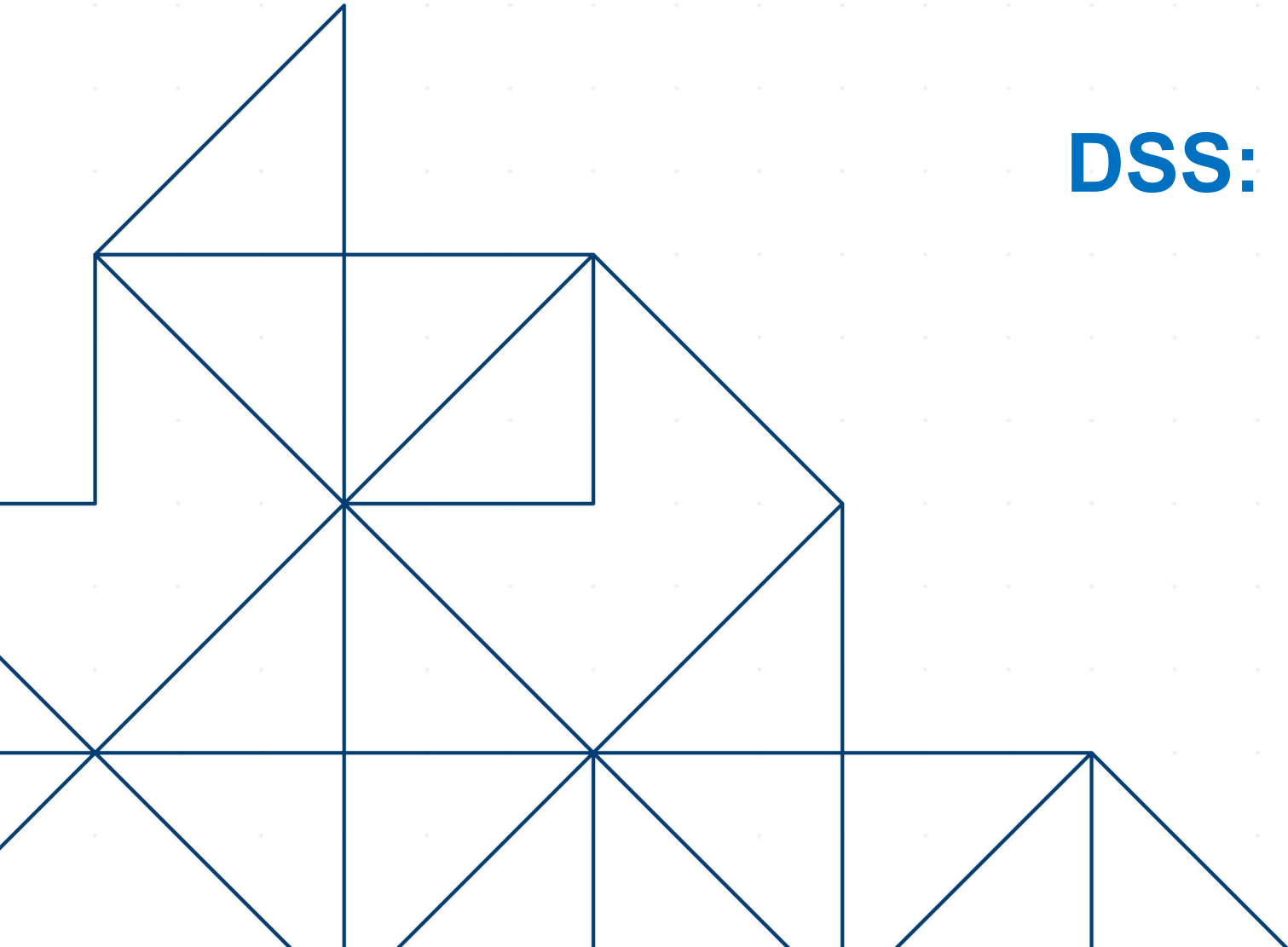
International Journal of
**Medical
Informatics**

Who is afraid of the system? Doctors' attitude towards diagnostic systems

Jacobus Ridderikhoff *, Bart van Herk

Erasmus University Rotterdam, Department of Family Medicine, PO Box 1738, 3000 DR Rotterdam, The Netherlands

Received 30 October 1997; accepted 21 June 1998



DSS: Intervene early



Research

Early diagnostic suggestions improve accuracy of GPs: a randomised controlled trial using computer-simulated patients

Olga Kostopoulou, Andrea Rosen, Thomas Round, Ellen Wright, Abdel Douiri and Brendan Delaney

British Journal of General Practice 2015; 65 (630): e49-e54. **DOI:** <https://doi.org/10.3399/bjgp15X683161>

N=198 GPs (UK)

N=100 GPs (Greece)

9 rich clinical scenarios

Range of difficulty

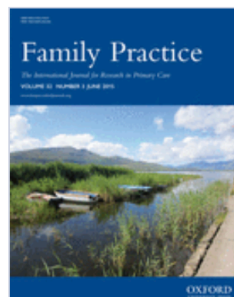
Online

Interactive

	Control	Early support
UK	63%	69%
GR	60%	71%

Pooled OR 1.40 [1.13, 1.67]

Family Practice

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Volume 32, Issue 3
June 2015

JOURNAL ARTICLE

Early diagnostic suggestions improve accuracy of family physicians: a randomized controlled trial in Greece

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[Olga Kostopoulou](#) ✉, [Christos Lionis](#), [Agapi Angelaki](#), [Salma Ayis](#), [Stevo Durbaba](#), [Brendan C Delaney](#)

Family Practice, Volume 32, Issue 3, June 2015, Pages 323–328,

Research

Diagnostic accuracy of GPs when using an early-intervention decision support system: a high-fidelity simulation

Olga Kostopoulou, Talya Porat, Derek Corrigan, Samhar Mahmoud and Brendan C Delaney

British Journal of General Practice 2017; 67 (656): e201-e208. **DOI:** <https://doi.org/10.3399/bjgp16X688417>

N=34 GPs

12 standardised patients (actors)

Within-participant design:

1. Baseline (6 patients): usual practice (EHR)
2. EHR+DSS (6 patients)
 - On different days
 - Patients counterbalanced

Research

Diagnostic accuracy of GPs when using an early-intervention decision support system: a high-fidelity simulation

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British Journal of General Practice 2017; 67 (656): e201-e208. DOI: <https://doi.org/10.3399/bjgp16X688417>

- Diagnostic accuracy: 8-9% absolute improvement (*OR* 1.41 [1.13, 1.77]; *OR* 1.50 [1.14, 1.99])
- Investigations: no significant increase
- Time: no significant increase

A recent clinical trial

Diagnoses supported by a computerised diagnostic decision support system versus conventional diagnoses in emergency patients (DDX-BRO): a multicentre, multiple-period, double-blind, cluster-randomised, crossover superiority trial

[Wolf E Hautz, MD](#) ^{a,*}  · [Thimo Marcin, PhD](#) ^{a,*} · [Stefanie C Hautz, PhD](#) ^a · [Stefan K Schaubert, PhD](#) ^b · [Prof Gert Krummrey, MD](#) ^{a,c} · [Martin Müller, MD](#) ^a · et al. [Show more](#)

- 4 emergency departments in Switzerland
- adults with fever of unknown origin, non-traumatic abdominal pain, syncope, or non-specific symptoms
- Isabel Pro DDx Generator, Isabel Healthcare, UK
- **No benefit**

Early enough?

Diagnoses supported by a computerised diagnostic decision support system versus conventional diagnoses in emergency patients (DDX-BRO): a multicentre, multiple-period, double-blind, cluster-randomised, crossover superiority trial

[Wolf E Hautz, MD](#) ^{a,*}  · [Thimo Marcin, PhD](#) ^{a,*} · [Stefanie C Hautz, PhD](#) ^a · [Stefan K Schaubert, PhD](#) ^b · [Prof Gert Krummrey, MD](#) ^{a,c} · [Martin Müller, MD](#) ^a · et al. [Show more](#)

- “Resident physicians inputted all signs and symptoms they deemed relevant into the prepopulated CDDSS, submitted those data, and received suggestions for possible differential diagnoses to consider from the CDDSS”
- Is consultation of a DSS immediately after taking a patient history and conducting a physical examination sufficiently early?


A single negative trial does not mean DSS is ineffective

THE LANCET
Digital Health

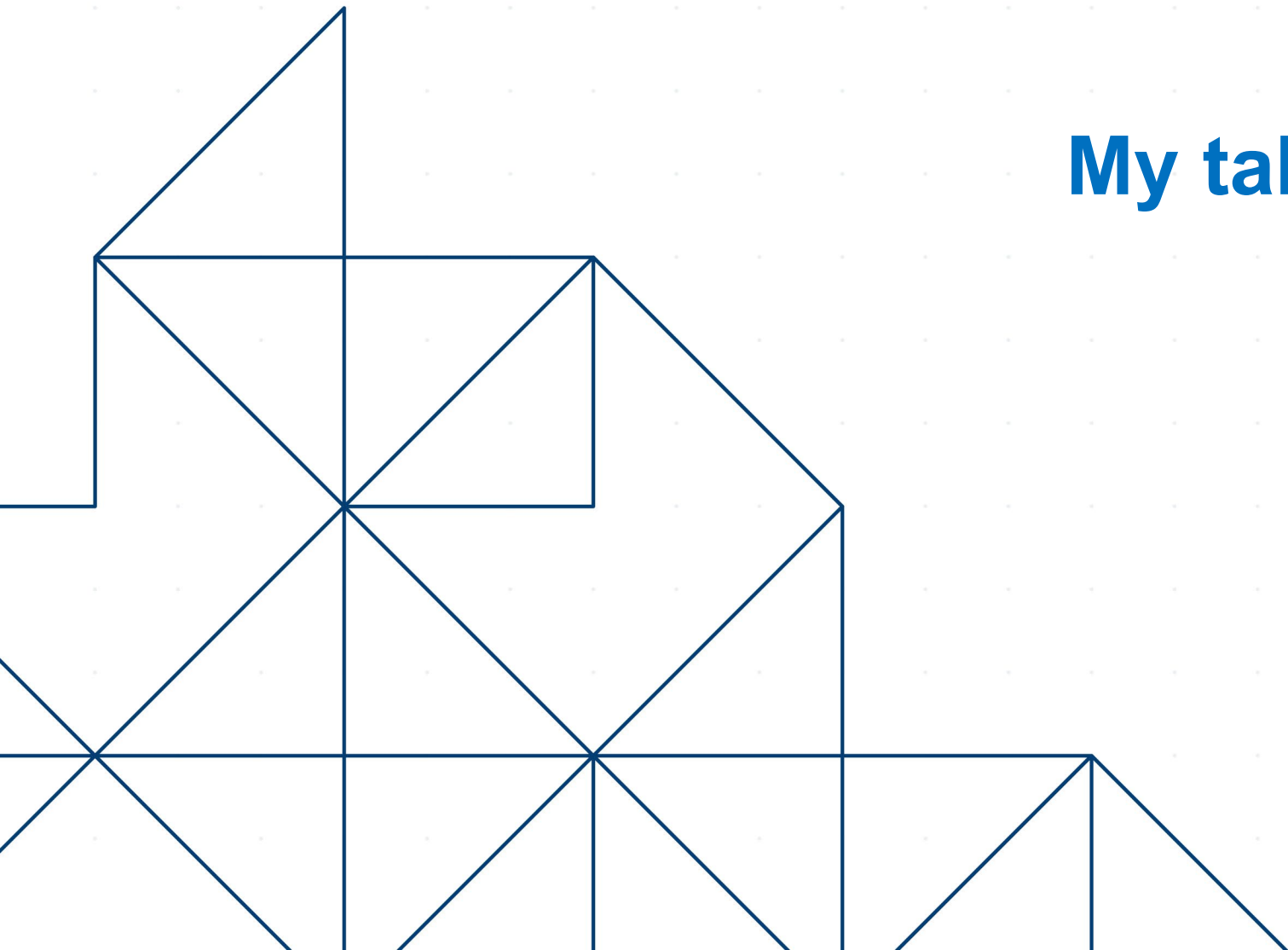
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COMMENT · [Volume 7, Issue 2](#), E108-E109, February 2025 · *Open Access* [Download Full Issue](#)

AI for medical diagnosis: does a single negative trial mean it is ineffective?

[Olga Kostopoulou](#)^a · [Brendan Delaney](#)^b 

- Implementation – need to understand exactly how AI tools are implemented and how they change cognition



My takeaway

Enabling good thinking strategies

Increasing AOT

ANNUAL REVIEW OF PSYCHOLOGY Volume 76, 2025

Review Article | Open Access

Boosting: Empowering Citizens with Behavioral Science

[Stefan M. Herzog](#)¹ and [Ralph Hertwig](#)¹

- Necessitates cognitive ability & motivation
- Self-nudging, habit formation, anticipating obstacles, changing the environment, reminders, apps & algorithms
- Long-term education and consistent training
- Safeguards autonomy and professional identity



AI: The Present & Future

- LLMs
- Explainable AI
- AI-powered, Ambient Voice Technologies (digital scribes)
- AI virtual doctors that collect data and suggest diagnoses before a clinician has met the patient.