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VOCAL AND SPEECH BIOMARKERS OF PSYCHIATRIC DISORDERS

Vincent P. MARTIN

February 1, 2023







HELLO!

I am Vincent P. MARTIN

- Ph.D. (2022) « Voice biomarkers of sleepiness »,
 Université de Bordeaux
 J.L. Rouas (LaBRI) & P. Philip (SANPSY/CHU)
- Eng. Degree (2018)
 Ecole Nationale Supérieure de l'Electronique et de ses Applications (ENSEA)
- DIU Philosophy of psychiatry (2021) Université de Bordeaux





I.
The need for objective diagnosis?

A NEED FOR OBJECTIVE DIAGNOSIS DEVICES?

"There is an **urgency** to **objectively diagnose**, monitor over time, and provide evidence-based interventions for individuals with mental illnesses"

Low et al. 2020

"Gold-standard diagnostic and assessment tools for depression and suicidality remain rooted, almost exclusively, on the **opinion of individual clinicians** risking a range of **subjective biases**. [...] Currently there is no **objective measure**, with **clinical utility**, for either depression or suicidality"

Cummins et al. 2015

A NEED FOR OBJECTIVE DIAGNOSIS DEVICES?

Aboraya 2007

- ▶ 28 healthcare professionals
- Most of them (87%): their diagnosis is not trustuworthy
- ▶ Why?
 - Factors linked to clinicians (education, biases, style): 63.5%
 - Patients characteristics: 21.6%
 - Disease definitions 14.9%

1rst factor for diagnosis
= clinician

A NEED FOR OBJECTIVE DIAGNOSIS DEVICES?

Kendell 1971

Table 6.—Diagnoses Given to Patient F						
	American Psychiatrists (N = 133)	British Psychiatrists (N = 194)				
Schizophrenia	92 (69%)	4 (2%)				
Simple	0	1				
Catatonic	1	0				
Paranoid	27	1				
Latent	8	0				
Residual	3	O				
Schizo-affective	33	1				
Unspecified	20	1				
Personality Disorder	10 (8%)	146 (75%)				
Paranoid	1	2				
Affective (cyclothymic)	1	8				
Explosive	0	2				
Hysterical	4	105				
Asthenic	0	2				
Antisocial	1	8				
Unspecified	3	19				
Affective Psychosis	10 (8%)	7 (4%)				
Neurosis	19 (14%)	37 (19%)				
Alcoholism or Drug Dependence	2	0				

⇒We need objective diagnosis

2. Voice biomarkers of psychiatric disorsers

State of the art

STATE OF THE ART

Low et al. 2020,
 « Automated assessment of psychiatric disorders using speech: A systematic review »,
 Laryngoscope Investigative Otolaryngology



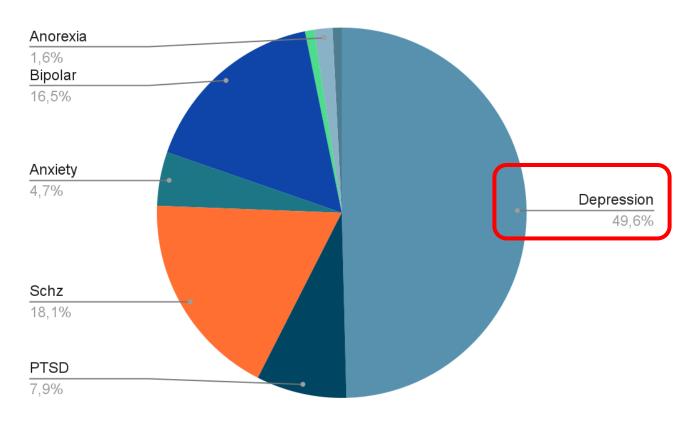
STATE OF THE ART Low et al. 2020 METHOD

- Google Scholar
- 2009-2019
- ▶ 127 studies



Supplementary data available online!

STATE OF THE ART Low et al. 2020 RESULTS



Pb: not used in real clinical settings

- Performances?
 - 80% for <u>bipolar disorders</u>, 95% for <u>schizophrenia</u>,
 89.3% for <u>depression</u>
 - High performances for decades (e.g. <u>75% for depression</u>)

- Performances? <a> \omega
- Databases size?
 - Ecological data collection
 - n=9920 (<u>Rutowski et al. 2022</u>)
 - n=3580 (<u>Di et al. 2021</u>)



- Performances? <a> \omega\$
- ▶ Databases ? ⁽²⁾
- Reglementary limitations?



WHY?

- Performances? <a> \omega
- ▶ Databases ? ②
- Reglementary limitations? <a> <a>
- Explanation and transparency?
 - TRUST



- Performances? <a> \omega\$
- Databases ? (3)
- Reglementary limitations? <a> <a>
- Explanation and transparency ? (3)



THERAPEUTIC RELATIONSHIP



Bourla et al.:

Bourla et al. 2018

- ▶ 515 psychiatrists
- ▶ 3 scenarios: biosensors comprising a connected wristband-based digital phenotype, ML-based blood test, ML-based magnetic resonance imaging (MRI).
- 4 acceptability domains usefulness, usability, reliability, and risk
- Overall acceptability=moderate.
- All systems = risky (410/515, 79.6%).
- Acceptability = strongly influenced by socioepidemiological variables (professional culture), such as gender, age, and theoretical approach.
- Worries = therapeutic relationship, data security, data storage, and privacy risk

THERAPEUTIC RELATIONSHIP

- Important for treatment adherence and therapeutic issues
- You may have depression », « You have a probability of XX% of having schizophrenia », …

3. Diagnostic: what are we talking about?

STATE OF THE ART Low et al. 2020 RESULTS

Label

Questionnaires (ex. PHQ9)



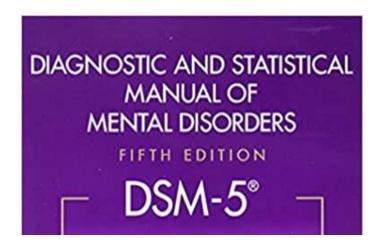
		Not at all	Several days	More than half the days	Nearly every day
1.	Little interest or pleasure in doing things	0	1	2	3
2.	Feeling down, depressed, or hopeless	0	1	2	3
3.	Trouble falling or staying asleep, or sleeping too much	0	1	2	3
4.	Feeling tired or having little energy	0	1	2	3
5.	Poor appetite or overeating	0	1	2	3
6.	Feeling bad about yourself — or that you are a failure or have let yourself or your family down	0	1	2	3
7.	Trouble concentrating on things, such as reading the newspaper or watching television	0	1	2	3
8.	Moving or speaking so slowly that other people could have noticed? Or the opposite — being so fidgety or restless that you have been moving around a lot more than usual	0	1	2	3
9.	Thoughts that you would be better off dead or of hurting yourself in some way	0	1	2	3

STATE OF THE ART Low et al. 2020 RESULTS

© Vincent P. Martin

Label

- Questionnaires (ex. PHQ9)
- Classification (e.g., DSM or ICD)



Major Depressive Disorder

Diagnostic Criteria

A. Five (or more) of the following symptoms have been present during the same 2-week period and represent a change from previous functioning; at least one of the symptoms is either (1) depressed mood or (2) loss of interest or pleasure.
Note: Do not include symptoms that are clearly attributable to another medical condition.

Depressed mood most of the day, nearly every day, as indicated by either subjective report (e.g., feels sad, empty, hopeless) or observation made by others (e.g.,

appears tearful). (Note: In children and adolescents, can be irritable mood.)2. Markedly diminished interest or pleasure in all, or almost all, activities most of the day, nearly every day (as indicated by either subjective account or observation).

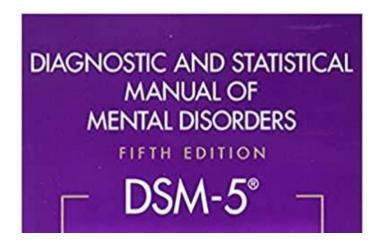
- 3. Significant weight loss when not dieting or weight gain (e.g., a change of more than 5% of body weight in a month), or decrease or increase in appetite nearly every day. (**Note:** In children, consider failure to make expected weight gain.)
- 4. Insomnia or hypersomnia nearly every day.
- Psychomotor agitation or retardation nearly every day (observable by others, not merely subjective feelings of restlessness or being slowed down).
 - 6. Fatigue or loss of energy nearly every day.
- Feelings of worthlessness or excessive or inappropriate guilt (which may be delusional) nearly every day (not merely self-reproach or guilt about being sick).
- Diminished ability to think or concentrate, or indecisiveness, nearly every day (either by subjective account or as observed by others).
- P. Recurrent thoughts of death (not just fear of dying), recurrent suicidal ideation without a specific plan, or a suicide attempt or a specific plan for committing suicide.

STATE OF THE ART (Low et al. 2020) RESULTS



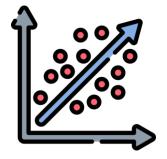
Label

- Questionnaires (ex. PHQ9)
- Classification (e.g., DSM or ICD)



Tasks

- diagnostic: binary classification
- severity estimation: regression with score



DIAGNOSIS: LIMITS

Questionnaires

- Not used by clinicians
- Validated on diagnosis criteria

Diagnosis criteria

- Culture (hikikomori)
- Time (versions of the DSM)
- Heterogeneity

DIAGNOSIS: LIMITS

Depression

- Number of semiological profiles
- = 326 unique profiles
- Eiko Fried: <u>STAR*D (2015)</u>: 1030 profiles on 3703 "depressive" patients

Major Depressive Disorder

Diagnostic Criteria

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- 2. Markedly diminished interest or pleasure in all, or almost all, activities most of the day, nearly every day (as indicated by either subjective account or observation).
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- Diminished ability to think or concentrate, or indecisiveness, nearly every day (either by subjective account or as observed by others).
- Recurrent thoughts of death (not just fear of dying), recurrent suicidal ideation without a specific plan, or a suicide attempt or a specific plan for committing suicide.

DIAGNOSIS: LIMITS

All pathologies Newson 2021

- 107349 patients
- ▶ **10** most prevalent disorders
- **47** symptoms

Conclusion: « DSM-5 disorder criteria do not separate individuals from random when the complete mental health symptom profile of an individual is considered.»

So, diagnosis == useless?



WHAT IS THE DIAGNOSIS USEFULL FOR?

All pathologies

- Acknowledgement from medical specialist and from society
- ▶ **10** most prevalent disorders
- 47 symptoms

- Conclusion:
 - « DSM-5 disorder criteria do not separate individuals from random when the complete mental health symptom profile of an individual is considered.»

WHAT IS THE DIAGNOSIS USEFULL FOR?

"the main aim of the psychiatric science is not classification as an end in itself but rather identification of causes and interventions"

Keneth Kendler, 2012

« [...] one of its most important goal is to facilitate communication among clinicians, researchers, administrators and patients [...] by establishing a common language." Derek Bolton, 2012

"[...] classification in itself is less important than often supposed to be, and less important than other tasks." Derek Bolton, 2012

- + prognostic
- + **differential** diagnosis

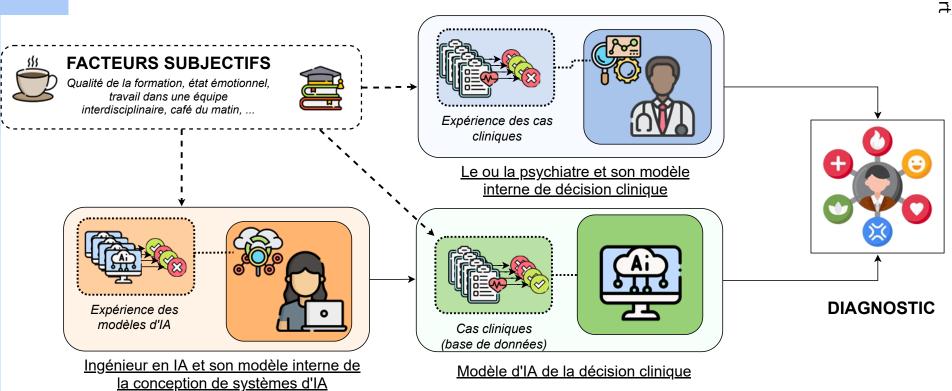
What is an w objective diagnosis »?

How Does Comparison With Artificial Intelligence Shed Light on the Way Clinicians Reason? A Cross-Talk Perspective

Vincent P. Martin^{1,2}, Jean-Luc Rouas¹, Pierre Philip^{2,3}, Pierre Fourneret⁴, Jean-Arthur Micoulaud-Franchi^{2,3} and Christophe Gauld^{4,5*}



'OBJECTIVITY' OF IA?



OBJECTIVITY: A GOLD-STANDARD?

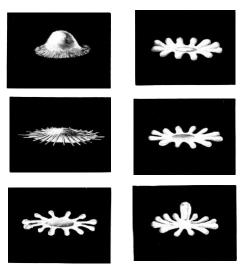
THE SPLASH OF A DROP, Pr. Worthington, Royal Institution of Great Britain, May 18, 1894

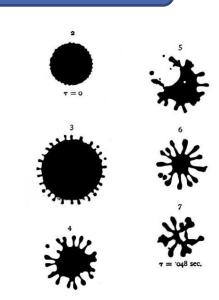
Mercury drop on glass

36

Light at constant delay

Which one represents **reality** more accurately?





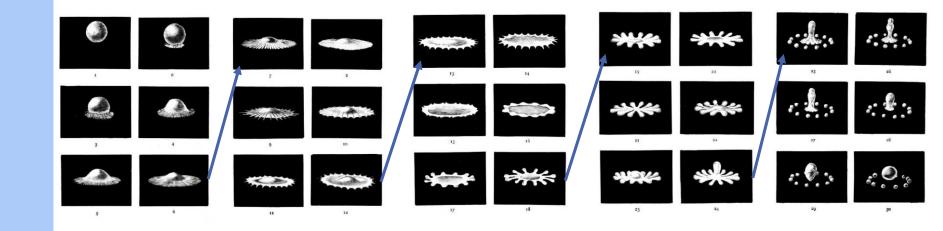
OBJECTIVITY: A GOLD-STANDARD?

THE SPLASH OF A DROP, Pr. Worthington, Royal Institution of Great Britain, May 18, 1894

Mercury drop on glass

37

Light at constant delay



4. What could be usefull for clinical practice?

Symptoms



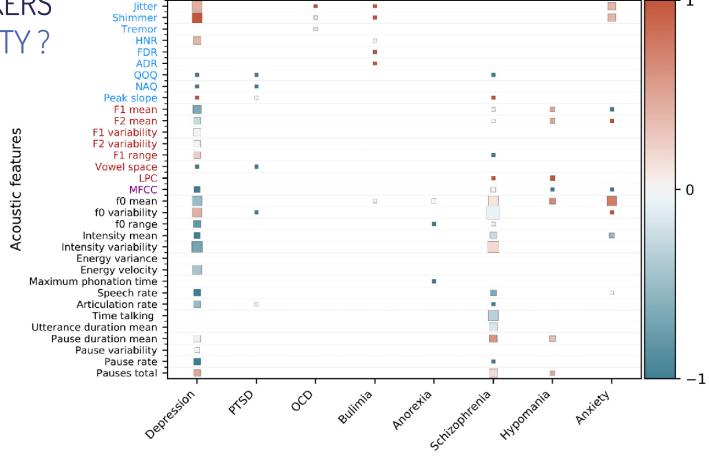
SYMPTOMS



Diagnosis	Symptoms
Time dependent e.g. DSM IV, DSM 5,	Stable through time
Cultural dependent e.g. Hikikomori	Independent from culture
Heterogeneous	Homogeneous
Symptoms → Syndromes → Diagnosis	
-	
-	







Psychiatric disorders

Low et al. 2020

SYMPTOMS



Diagnosis	Symptoms
Time dependent e.g. DSM IV, DSM 5,	Stable through time
Cultural dependent e.g. Hikikomori	Independent from culture
Heterogeneous	Homogeneous
Symptoms → Syndromes → Diagnosis	
-	Mechanistic explanation
-	



SYMPTOMS



Diagnosis	Symptoms
Time dependent e.g. DSM IV, DSM 5,	Stable through time
Cultural dependent e.g. Hikikomori	Independent from culture
Heterogeneous	Homogeneous
Symptoms → Syndromes → Diagnosis	
-	Mechanistic explanation
-	Necessary for diff. diag and prog.



"subjects have no need to be equipped with multiple sensors or even be burdened by **invasive devices** (e.g., endoscopy) [...] Additionally, CA can make it feasible to collect data from subjects via mobile devices (e.g., a smartphone), which can provide the subjects **24×7 monitoring service**." *Qian et al. 2020*



- + no judgment
- + easy to access (smartphones)
- + no biases (patients and clinicians

Anchoring bias	Tendency to focus on a first impression or on the first information received to form an opinion about a number, a person, an event This judgmental
	blas can prevent important information received later to be taken
	into account.
Ascertainment	Tendency to selectively analyse clinical data in the light of prior expectations or beliefs (belief bias). This bias can impact the interpretation of new
	information resulting from precise surveillance or screening of
	certain symptoms.
Availability bias	Tendency to form an opinion based on the most recent and readily available information in one's mind, considered more likely. For example, for an opinion on a treatment, we remember the last few patients rather than a series of 100.
Base-rate neglect	Type of error due to poor knowledge of disease incidence rates, either by
	underestimating or by overestimating the occurrence of a diagnosis.
Confirmation bias	Tendency to select and interpret information confirming a clinical intuition or a priori diagnosis, and to neglect information that contradicts or invalidates this intuition.
Diagnosis momentum	Diagnosis or treatment plans established by previous clinicians are rarely
	questioned by new practitioners and stick to the patient. This phenomenon can prevent considering new options and enhancing the diagnosis or provided healthcare.
Illusory correlation	Tendency to infer causation relationships between correlated but independent events.
Premature closure	Tendency to stop reasoning, evaluating or looking for a better diagnosis or treatment alternative after finding a suitable enough option (close to 'satisfaction search bias').
Primacy effect	Mnemonic bias, tendency to remember and consider more the first information out of a list of equal importance.
Recency effect	Mnemonic bias, tendency to remember and consider more the most recent information (received last), for example the last words of a clinical interview or the last symptoms of a list.
Unpacking principle bias	Type of error occurring when not all the necessary information were requested to make an objective judgement. The risk would be, for example, to omit
Affect bias	information that would allow a differential diagnosis. When decisions are made in a context where the immediate emotions are
Affect bias	when decisions are made in a context where the immediate emotions are strong and can influence our choices.
Ambiguity or risk aversion	Type of bias describing the tendency to favour choices with known risks and
randigately of the decision	associated probabilities rather than ambiguous or uncertain options.
Commission bias	Tendency to favour action over inaction, even when inaction would be more rational. It can result in overprescription.
Default bias or status quo bias	Tendency to stick to the default option and avoid changes. The cost of change
	in terms of cognitive effort is automatically considered too great and one continues to behave in the same way.
Framing bias	The perception of a situation can be influenced by the way options are being presented (formulation with different numerical presentations, or with positive or neadtive connotations).
Information bias	This bias translates into errors in the collection of information, for example
	during an interview: it can be a failure to observe, a misclassification or organisation of data, or errors in memory recall during synthesis.
Loss aversion	Tendency to be more sensitive to the loss of a certain amount of resources
	(cognitive effort, time, money) than to the gain of the same amount of resources, resulting in choices that tend to avoid losses rather than
Omission bias	attempt gain. Tendency to favour inaction or to avoid difficult issues over action ('wait and
	see'). It affects self-doubting clinicians.
Outcome bias	Tendency to focus on the outcome of the decision rather than the information
	to be interpreted to make a relevant decision. This bias is more common among clinicians with lower self-confidence and can lead to an

Tendency to rely on the 'frequency argument,' i.e., to favour the most common hypotheses and not to mention the rarer ones. It is a restriction of thought that prevents a broader questioning of a clinical situation. When the result of a situation is known, it can influence the way in which we perceive the preceding events as we forget the uncertainty we were facing at that time, and lead to fallacious reconstruction ('we are remaking history'). It can prevent learning and lead to the repetition of error. Tendency to reduce the analysis of clinical data and the diagnosis to one's own point of view. It affects communication between the different parties (physician, patients, and other stakeholders). Tendency, when one has already invested a lot of resources (time, energy or money) in a project or an action that seems to have little chance of succeeding, to continue investing although it is doomed to failure. In medicine, it is a question of pursuing an ineffective strategy, for example. Tendency to conform and reproduce a behaviour or an attitude just to act as others do. While making judgments about people's behaviour, it's the tendency to overemphasise dispositional factors or personality-based explanations and underestimate situational ones. The consequence is the risk of making incorrect judgments, discounting reasons that might have contributed to their observed behaviour. Tendency to infer characteristics about an individual based on the group in which we categorised him/her. This can result in a wrong diagnosis solely

based on our belief that the patient belongs to a certain group with a

Mouchabac et al. 2021

typical disease.

Representativeness restraint bias

Retrospective prejudice

Self-served bias

Sunk cost fallacy

Bandwagon effect

Stereotyping

Fundamental attribution error

"subjects have no need to be equipped with multiple sensors or even be burdened by **invasive devices** (e.g., endoscopy) [...] Additionally, CA can make it feasible to collect data from subjects via mobile devices (e.g., a smartphone), which can provide the subjects **24×7 monitoring service**." *Qian et al. 2020*



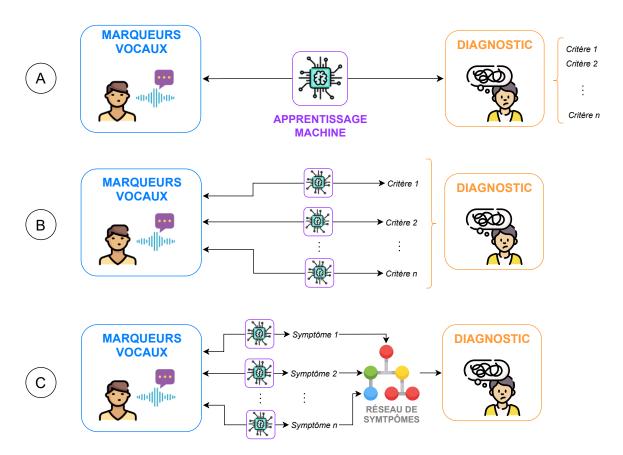
- + no judgment
- + easy to access (smartphones)
- + no biases (patients and clinicians)

"subjects have no need to be equipped with multiple sensors or even be burdened by **invasive devices** (e.g., endoscopy) [...] Additionally, CA can make it feasible to collect data from subjects via mobile devices (e.g., a smartphone), which can provide the subjects **24×7 monitoring service**." *Qian et al. 2020*

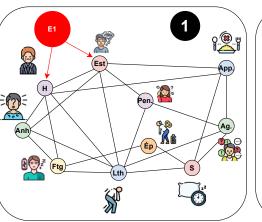


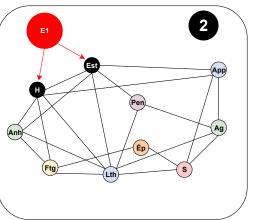
- + no judgment
- + easy to access (smartphones)
- + no biases (patients and clinicians)
- epistemic injustices

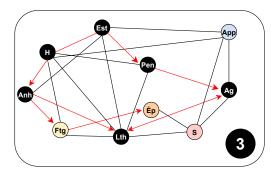
SYMPTOMS NETWORKS

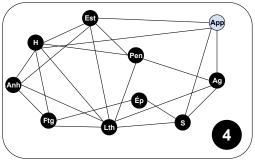


SYMPTOMS NETWORKS

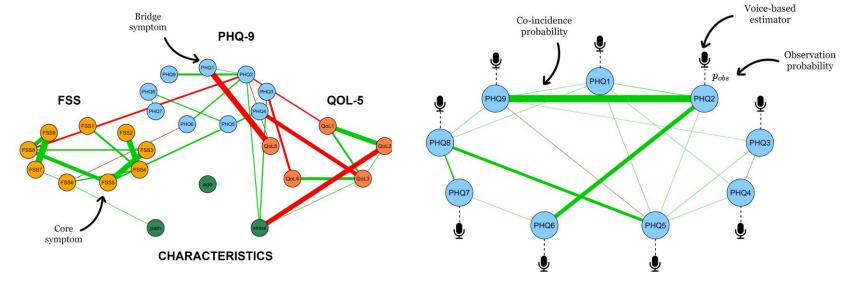








SYMPTOMS NETWORKS



Conclusion

Doggy bag

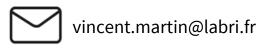
DOGGY BAG

- Beware of preconceived ideas about medical practice when you are a computer scientist (get trained!)
- Objectivity is not necessarily better than subjectivity
- Symptoms instead of diagnoses

Thank you for your attention!



QUESTIONS?

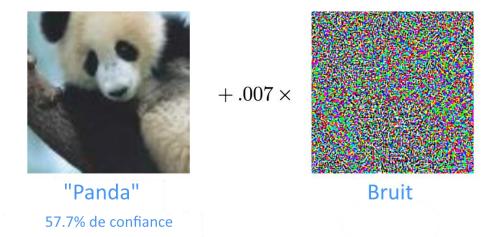






BONUS: «BUT IT WORKS!»

Adversarial attacks



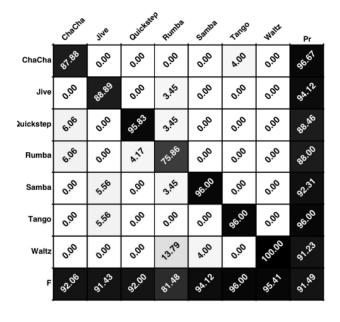
- Adversarial attacks
- Digression: music information retrieval

<u>Sturm 2016</u>

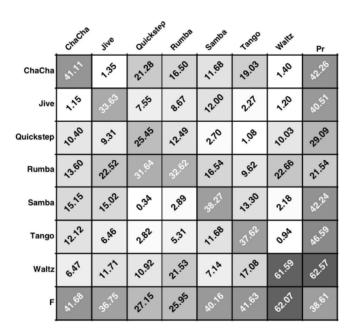
2013 MIREX challenge « Audio Latin Music Genre classification task »



BONUS: « BUT IT WORKS!»



Time stretching 32 values in [0.85, 1.15]



- Adversarial attacks
- Digression: music information retrieval
- ▶ The horse of Dr. Von Osten (Pfungst,1911)



- Adversarial attacks
- Digression: music information retrieval
- ▶ The horse of Dr. Von Osten (Pfungst,1911)

Solution?

Irrelevant transformations



THE END

DEEP LEARNING?

- IS19 challenge: winner = Fischers vectors + SVR Recent DL models: perf < IS19
- 2. C. Rudin 2019 « Stop explaining black box machine learning models for high stakes decisions and use interpretable models instead », *Nature Machine Intelligence*
- 3. Meta analyses: Christodoulou « A systematic review shows no performance benefit of machine learning over logistic regression for clinical prediction models» *J. Clinic. Epidemio*.
- → Did you put as much efforts in logistic regression than in tunning a deep learning model?

IS EXPLAINABILITY ENOUGH?

Vilone et al. 2021 "Notions of explainability and evaluation approaches for explainable artificial intelligence", Information Fusion

Algorithmic The degree of confidence of a learning algorithm to behave 'sensibly' in general [2],

transparency

[26]

Causality The capacity of a method for explainability to clarify the relationship between input

and output [8], [21], [22], [23], [24], [25], [29]

Comprehensibility The quality of the language used by a method for explainability [9], [31], [32], [33], [34],

[35], [36], [37], [38]

Effectiveness The capacity of a method for explainability to support good user decision-making

[40], [41], [42], [43]

Efficiency The capacity of a method for explainability to support faster user decision-making

[20], [41], [42]

Explicability The degree of association between the expected behaviour of a robot to achieve

assigned tasks or goals and its actual observed actions [44]

Explicitness The capacity of a method to provide immediate and understandable explanations [45]

Faithfulness The capacity of a method for explainability to select truly relevant features [45] Interestingness The capacity of a method for explainability to facilitate the discovery of novel

knowledge and to engage user's attention [33], [34], [36], [53], [54]

Interpretability The capacity to provide or bring out the meaning of an abstract concept [9], [18], [33],

[35], [55], [56], [57], [58], [59], [60], [61], [62], [63]

The capacity of a method for explainability to provide useful information to end-Informativeness

users [21]

Justifiability The capacity of an expert to assess if a model is in line with the domain knowledge [1],

[33], [40], [55], [64], [65]

Mental Fit The ability for a human to grasp and evaluate a model [33], [66]

The capacity of a method for explainability to convince users perform certain actions Persuasiveness

[20], [41], [42]

Selection/ The ability of a method for explainability to select only the causes that are necessary

and sufficient to explain the prediction of an underlying model [25]

The extent to which each component of an explanation's content is truthful in Soundness

describing an underlying system [27], [28]

The capacity of a method to explain how the system works even when it behaves Transparency

unexpectedly [9], [10], [11], [12], [20], [26], [40], [41], [47], [58], [59], [63], [64], [76], [77], [78]

Structure of an explanation

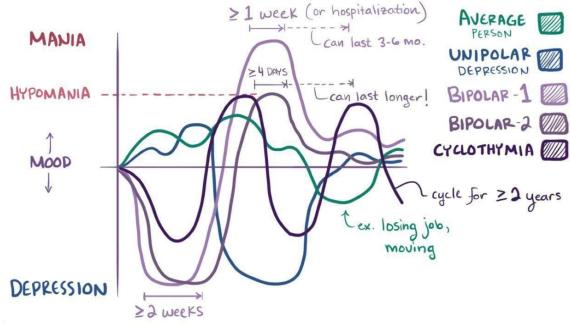
simplicity

End-user Goals Contents Language What questions What information Which Who is the receiver of the explanation? should be should be terminology answered? contained in it? should be used?

WHAT DO WE DETECT? EXAMPLE 1: BIPOLAR DISORDER

Bipolar disorderS

- Diag. = based on variations and duration
- How to detect BD with only 1 recording?
- State vs. Trait (same for all Ψ disorders)



https://www.osmosis.org/learn/Bipolar_disorder

WHAT DO WE DETECT? EXAMPLE2: DEPRESSION

What does a ML classifier learn?

- Difference between groups
 - Sub-group?
 - Symptom?
 - Other bias?
 - **>** 3
- + /!\ Temporality /!\

Depressive vs. HC with bad mood

NOT "depression disorder"

