

### What you need to know about Delirium in ICU

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#### **Delirium and outcome**

40 year old ARDS ICU survivor college graduate "I have been out of hospital and trying to get on

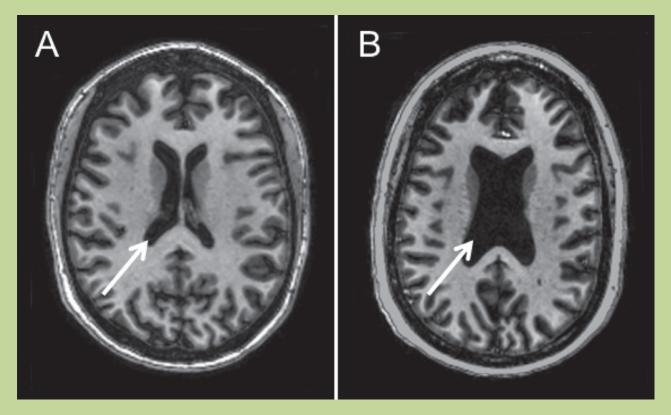
with my life for the past 2 years. I have trouble with people's names that I have worked with for years. I can't remember where I put things at home. I can't help my children with their homework because I can't remember how to do simple multiplication problems."

## What is happening in the brain?

- Oxidative stress
- Neurotransmitter imbalance
- Neuronal aging
- Inflammation
- Abnormal levels of large neutral amino acids



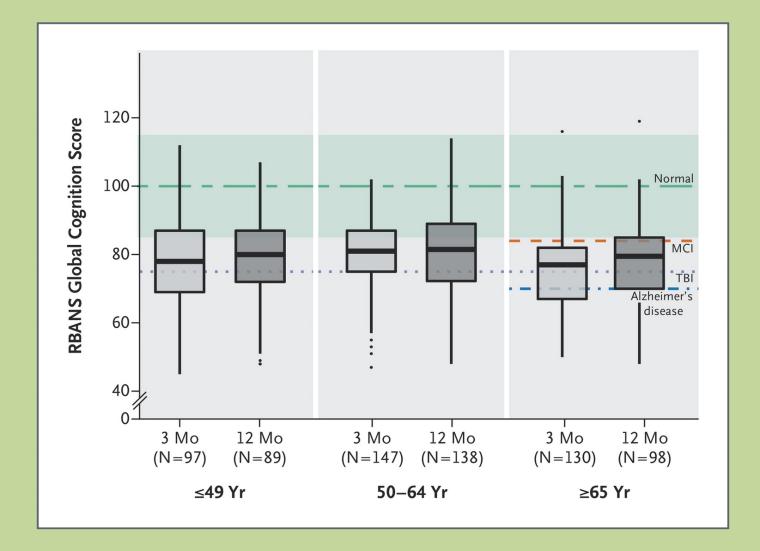
#### VISIONS



A 46 year old woman ARDS, no delirium. B 42 year old woman ARDS, 12 days delirium <sub>Gunther CCM 2012</sub>



#### **Global Cognition Scores in Survivors of Critical Illness.**



Pandharipande P et al. N Engl J Med 2013;369:1306-1316.



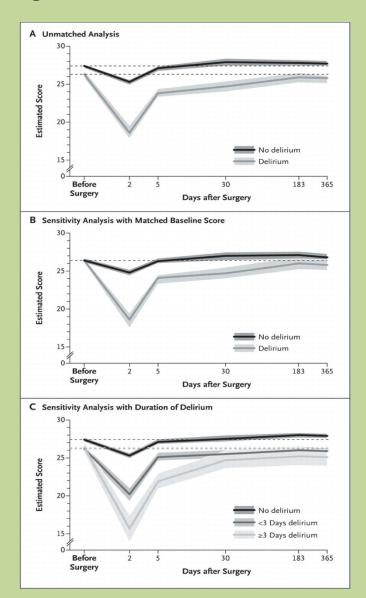
### Post-operative cardiac surgery

- 225 patients 60 years or over
- CABG or valve replacement
- Cognitive function monitored with MMSE
- Delirium diagnosed using CAM
- KATZ index of ADLs



#### **Results** -trajectories

46% patients developed delirium Time to return to preoperative MMSE No delirium: 1 month. Delirium: not returned by 1 year, still improving up to 6 months



#### **Cognitive outcomes Hope-ICU** Telephone Interview of Cognitive Status

	Haloperidol	Placebo
TICS-M Median (IQR) n = 57	22 (18-27)	21 (18-24)

≤ 31 cutoff score separates MCI from normal cognition (sensitivity 71.4%)

≤ 27 score separates MCI from dementia (sensitivity 69%).

#### The delirium experience

"The rest of my stay in ICU was filled with more incidents of despair, humiliation and terror. I saw a patient stabbed to death by his wife, and two people committing suicide. I witnessed arguments, in my mind all caused by me, and the pain I felt as my lungs started to recover was all part of a plan to give me pain inducing drugs – in fact I had seen doctors laughing about it.

The day after I was extubated I found myself in the High Dependency Unit, where the sheer terror of the execution attempts began."

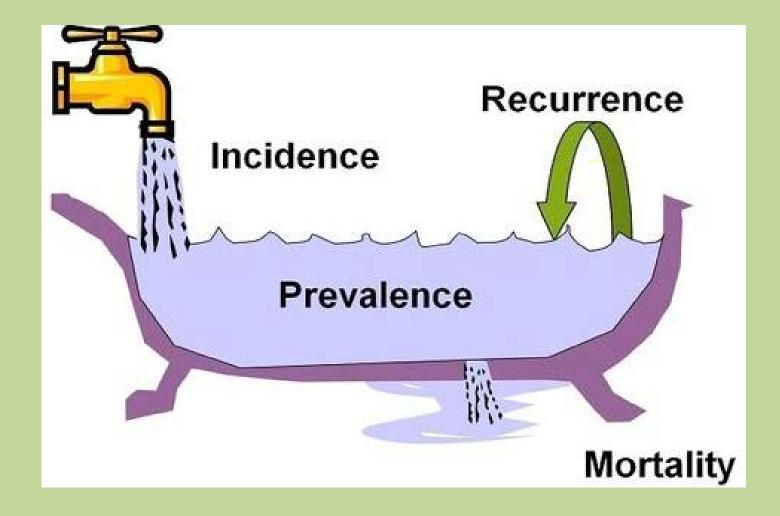
#### Delirium – DSM V

- Disturbance in attention and awareness
- Acute onset and fluctuates
- Disturbance in cognition
- Not explained by pre-existing, established or evolving neurocognitive disorder and noncomatose patient
- Evidence for cause

American Psychiatric Association. (2013). *Diagnostic and statistical manual of mental disorders* (5th ed.). Arlington, VA: American Psychiatric Publishing.

#### Subsyndromal delirium

- One or more symptoms
- Not meeting full criteria
- Not progressing to delirium
- Intermediate outcomes



#### North America

- SLEAP trial 420 ventilated patients
  - APACHE 23
  - 54% delirium,
  - 4% ICDSC 0 throughout
- BRAIN trial 821 patients
  - APACHE 25
  - 74% delirium
- Mehta S, Cook D, Devlin JW, et al Prevalence, risk factors, and outcomes of delirium in mechanically ventilated adults. *Crit Care Med.* 2015; **43** :557-66.
- P.P. Pandharipande, T.D. Girard, J.C. Jackson, et al. Long-Term Cognitive Impairment after Critical Illness. *N Engl J Med* 2013; **369**:1306-16.

#### Netherlands

Netherlands:

Before/after study: 476 patients before/after study 44% to 97%

## Observational study: 1112 consecutive patients 50% at least 1 episode

van den Boogaard M et al. Haloperidol prophylaxis in critically ill patients with a high risk for delirium. *Crit Care.* 2013 Jan 17;**17**:R9.

Klowenberg K et al. The attributable mortality of delirium in critically ill patients: prospective cohort study. *BMJ* 2014; 349: g6652

#### World ICU population

- Belgium 15%
- Germany 26.9%
- Sweden 39%
- Australia 12.4%
- UK 32.4%

Van den Boogaard Intensive care medicine 2014; 40: 361-9

#### Incidence

Acute general hospital 1000 beds = 100 delirious patients

#### **Critical Care**

- 1:5 high dependency up to 4:5 sick ventilated patients
- Compare 3:5 with fractured neck of femur

## Watford Incidence of delirium by subgroup

Crit Care 2009; 13: R16

	Delirious	Not delirious
Elective post-operative (n=23)	1 (4%)	22 (96%)
Emergency admissions (n=57)	22 (45%)	27 (55%)
Ventilated patients (n=27)	17 (63%)	10 (37%)

#### What does it look like?







### **Delirium motoric types**

- Hyperactive psychomotor agitation
- Hypoactive psychomotor lethargy and sedation, appears quiet & co-operative BUT with inattention and disorganised thinking.
- Mixed fluctuating hypo/hyperactive symptoms

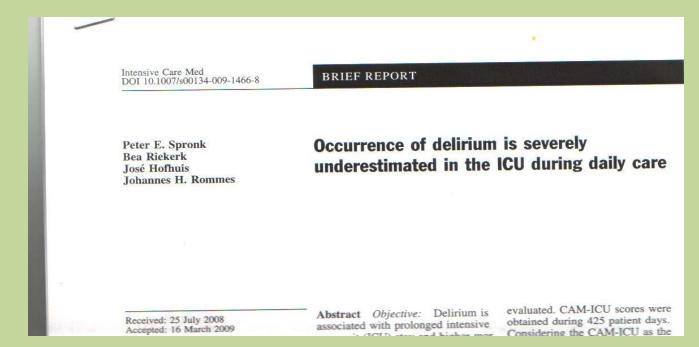
#### Identification



## CAM-ICU



#### **Diagnosing delirium**



46 patients Median age 73 years (IQR = 64–80) CAM-ICU scores during 425 patient days Delirium in 50% of the patients 3 days (range 1–9). Clinical detection 24%

#### Rapidly Reversible, Sedation-related Delirium versus Persistent Delirium in the Intensive Care Unit

Shruti B. Patel, Jason T. Poston, Anne Pohlman, Jesse B. Hall, and John P. Kress

Department of Medicine, Section of Pulmonary and Critical Care, University of Chicago, Chicago, Illinois

# Can delirium be diagnosed in sedated patients?

Patel SB. AJRCCM 2014;189:658-65 Takala J. AJRCCM 2014;189:622-24

#### Sedation related delirium

- 102 of 256 patients
- Paired CAM-ICU before and after SAT
- 28.9% CAM-ICU (n=11)negative after SAT
- 89% at least 1 day delirium pre vs. 77% post.
- Outcomes, same for rapidly reversible as no delirium.

Can delirium be diagnosed in comatose patients?

#### We Know

- •More sedation, more delirium
- •More delirium, worse outcomes
- •Less sedation, better outcomes

Balas MC et al Crit Care Med 2014, Jackson DL et al Crit Care 2010, Dale CR Ann Am Thorac Soc 2014, Shehabi Y et al Intensive Care Med 2013, Tanaka LMS et al, Crit Care 2014, Porhomayon J et al. J Cardiovasc Thorac Res 2015, Girard TD et al. Lancet 2008; 371 (9607) 126-134

#### Sedation induced coma RASS -3 to -5

- SPICE/ANZICS trial deep sedation throughout the first 48 hours in 171 (68%) patients
- SPICE group Malaysia deep sedation in 159 (61%) patients and 1,658 (59%) of all RASS assessments at 48 hours.
- ABC trial day 1, 52% control group, 56% intervention group were in coma.

Shehabi Y, et al. Intensive Care Med 2013; 39: 910-8,

Shehabi Y et al Sedation Practice in Intensive Care Evaluation (SPICE) Study Investigators; ANZICS Clinical Trials Group. Am J Respir Crit Care Med. 2012; **186**: 724–31

Girard TD et al. Lancet 2008; 371: 126-34

#### Watford – first 2 days

- Average number of RASS per shift 3.56
- Number of RASS -1 to +1 = 18.7%
- In 28.3% of patients
- No correlation with APACHE



### **Delirium or Depression?**

- 'The symptoms of delirium, when they are present, are prominent and take diagnostic precedence as delirium frequently indicates serious and urgent physical pathology or morbidity.'
- 'Because of their clinical overlap, in a hospital setting, clinicians should not rule out delirium in any patient with suspected depression – '

Take home messages: O'Sullivan et al Lancet Psychiatry 2014

#### **Delirium or Depression?**

#### **Overlapping features**

- Affective changes
- Sleep disturbance
- Underactivity or lethargy
- Apathy
- Agitation
- Impaired speed of information processing
- Delusions or hallucinations
- Impaired memory

#### Delirium – Prominent features

- Fluctuation in intensity of symptoms
- Acute onset
- Altered consciousness
- Marked inattention
- Underlying physical cause
- Disorientation
- Disorganised thinking
- Poor comprehension

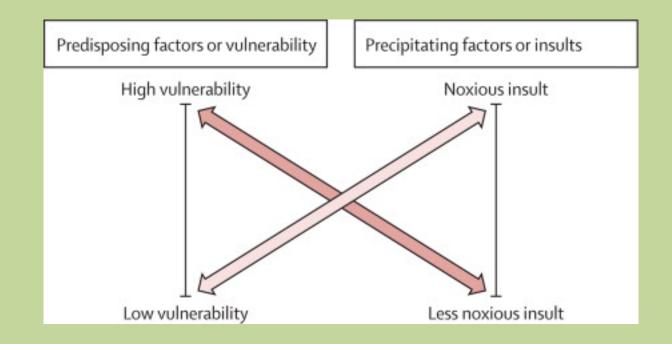


Figure. Multifactorial model of delirium in older people

Onset of delirium is dependent on a complex interaction between the patient's baseline vulnerability (predisposing factors) at admission and precipitating factors or noxious insults.

Sharon K Inouye, Rudi GJ Westendorp, Jane S Saczynski **Delirium in elderly people** 

, Volume 383, Issue 9920, 2014, 911-922

#### 

#### **Risk factors**

Host factors	Acute illness	Iatro/environ	
Elderly	Severe sepsis	Sedative/analges	
Co-morbidities	ARDS	Immobilisation	
Pre-existing	MODS	TPN	
cognitive impair			
Hearing/vision	Drug OD or	Sleep	
impairment	illicit drugs	deprivation	
Neurological dis	Nosocomial inf.	Malnutrition	
Alcohol/smoker	Met. disturbance	Anaemia	

#### Management

- Sedation score and delirium screening
- <u>Identify and treat</u> <u>precipitating factor</u>
- Minimise impact of predisposing factors
- Pharmacological therapy

Specimen type Blood Sodium 140 mmol/L (135 - 14	eGFR	30 L mL/min	
Potassium 4.8 H mmol/L (3.2 - 4.   Chloride 110 mmol/L (100 - 11)   Bicarb. 23 mmol/L (22 - 33)   Anion Gap 9 mmol/L (4 - 13)   OSM(Calc) 295 mmol/L (3.0 - 7.   Glucose 9.2 H mmol/L (3.0 - 7.	.5) Protein 10) Albumin 0 Globulin Bilirubin 95) CK 8) ALP 0) Gamma GT 0) ALT 1) AST	1.73m <sup>2</sup> 44 L g/L 23 L g/L 21 L g/L 70 C umol/L 215 H U/L 98 U/L 16 U/L 125 H U/L 215 H U/L 215 H U/L 2690 H U/L	2 (62 - 83) (33 - 47) (25 - 45) (< 20) (< 20) (40 - 110) (< 50) (< 45) (< 40) (110 - 250)
Diff: Manual Specimen Hgb : 83 L WBC : 11.4 PLT : 44 L : RBC : 2.72 L HCT : 0.24	4 H		
GENERAL COAGULATION INR 1.2 Prothrombin Time 12 APTT 27 Fib (derived) 8.0	н		

#### Predisposing factors? Management – non-pharmacological

"Delirium bundle", optimisation of risk factors

- Address visual, hearing impairment
- Orientation
- Familiar nurse
- Mobilisation
- Drug overhaul
- Sleep

Naughton et al. J Am Geriatri Soc 2005;53:18-23, Lundstrom et al J Am Geriatri Soc 2005;53:622-28

#### Drugs and delirium

- Opiate analgesics
- Benzodiazepines
- Corticosteroids
- Anticholinergic load Furosemide Ranitidine Digoxin

# Brain road map for rounds (script for interdisciplinary communication)

# Skipping any of these steps could leave the clinical team wanting more information!

Investigate (Ask these questions)	Report (only takes 10 seconds)
Where is the patient going?	Target level of consciousness (RASS, SAS)
Where is the patient now?	Actual level of consciousness (RASS, SAS) Delirium assessment (CAM-ICU, ICDSC)
How did they get there?	Drug exposures

#### SLEEP, ITS IMPORTANCE IN PREVENTING INSANITY

We wish we could impress upon all, the vast importance of securing sound and abundant sleep ; if so, we should feel that we had done an immense good to our fellow-beings, not merely in preventing insanity, but other diseases also.

American Journal of Insanity Boston Med Surg J 1845; 32:299-301

### **Deep sedation**

### **Sleep deprivation**

#### Sleep and sedatives

- Not cyclical medication specific, dose dependent
- Decrease in SWS & REM sleep
- ?restorative
- Benzos and Propofol activate GABA in VLPO suppress histamine release from TMN
- NA release from LC not inhibited
- Overdose burst suppression

### Sleep disturbance - sedatives

- Observational study 21 ventilated patients
- Polysomnography
- Opiates 100%, Benzo 23%, Propofol 82%
- Normal features of sleep not identified
- REM in 2 subjects
- ?Altered state of consciousness, continual sleep deprivation

Gehlbach et al Sleep 2012; 35: 1105

#### Delirium

- Does sleep deprivation contribute to delirium?
- Does delirium contribute to sleep deprivation?
- Review of 17 surgical studies not a risk factor
- Plausible but not proven

#### Case story

- Independent
- Lives alone
- Widow 2 years
- Was refused hip operation
- Suffers from pain

### Day 8/9

- RASS +4
- Intermittent fast AF
- Palliative care referral.
- Continued quetiapine at night
- 0.5mgs lorazepam iv
- Lines out
- Continue NG feed

### Day 10

- Woke up
- Asked for her glasses
- RASS 0
- CAM-ICU positive
- Delirium cleared
- Discharged to ward

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# Patients

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#### Welcome to your preview of The Times

#### My nightmare in hospital

David Aaronovitch Last updated at 5:45PM, November 12 2011

David Aaronovitch explains how routine keyhole surgery led to a terrifying bout of 'ICU psychosis' during which he thought he was going mad

On Sunday, September 4, I woke up to find that I was no longer mad. It was 2pm, my two brothers were sitting on either side of my hospital bed, my wife between them, the sun was slanting in through the window behind me and the horror that had dominated my life for nearly



David Aaronovitch, photographed last month Mark Harrison Print Email Share

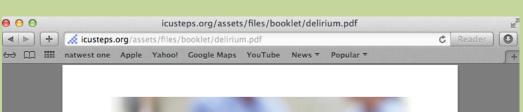
Login Subsci

Thurs

a week had evaporated. But I will never forget those days and nights of terror and delusion, and will never think about madness in the same way again.

#### **Friends and Family**

#### www.icusteps.org





#### What is delirium?

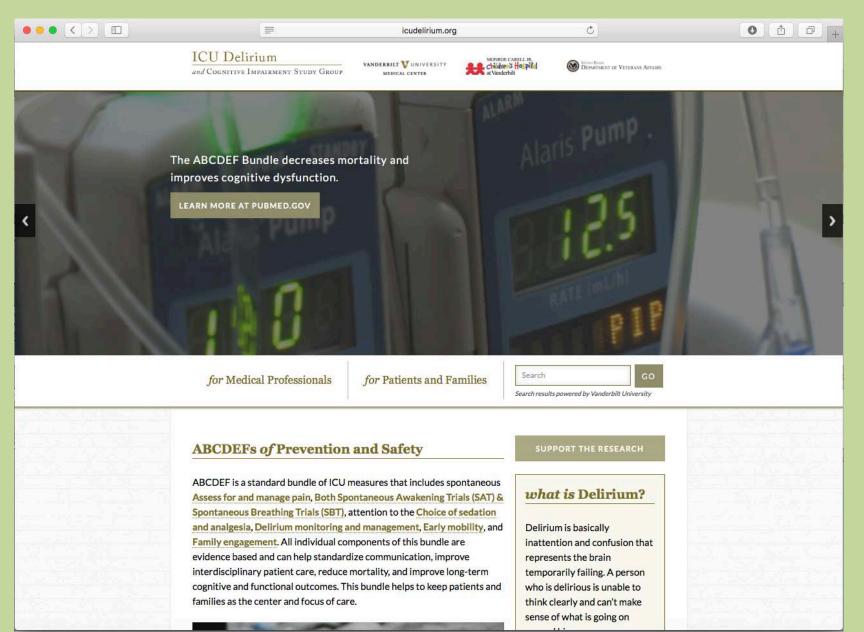
Delirium is a name for acute confusion. The patient who is delirious is often experiencing a world that makes no sense to us but is very real to them. For instance they may:

- not know they are in hospital
- think they can see frightening animals
- think they have been kidnapped
- think staff are only pretending to be nurses
- try to make sense of the noises around them and create a different explanation for them, so for instance if another patient is upset, they may think someone is being tortured.

The main point is that the patient is absolutely convinced about the reality of the confused world they are in. It can be terrifying for them and very worrying for relatives.

Often a patient who is delirious will still recognise friends and family although they will not generally believe their reassurances. They will usually want to get out of bed and be taken home. Patients with delirium can find it very difficult to understand or retain information – so even if they appear to understand what is happening, or may be joining in a conversation, they may not remember what has just been said to them. Delirium can also fluctuate, one minute you will be having a normal conversation and next they will say something that makes no sense.

#### www.icudelirium.org



#### Acknowledgements

- Professor Danny McAuley, Queens University, Belfast
- Drs Tim Alce & Annalisa Casarin Research Fellows
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- Dr Neil Soni, Imperial College, London
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