Acute Encephalopathy in a Hospitalized Patient

Acute encephalopathy is an altered mental status (AMS) due to diffuse brain dysfunction, often reversible with prompt evaluation and management.

Definition and Epidemiology

- **Definition:** Acute encephalopathy is a rapid-onset (hours to days) change in mental status (confusion, delirium, coma) due to systemic or neurologic insult, often reversible.
- Prevalence: ~20% of hospitalized patients; 50-70% in ICU (delirium, sepsis).
- Risk Factors: Age >65, ICU stay, sepsis, polypharmacy, substance use (alcohol, opioids), prior neurologic or psychiatric disease.

Pathophysiology

- Mechanisms: Diffuse brain dysfunction from metabolic, toxic, infectious, structural, autoimmune, or psychiatric causes → Impaired neuronal signaling.
- Effects: Altered consciousness (confusion to coma), cognitive decline, seizures.
- Key Pathway: Systemic inflammation (e.g., sepsis) or immune dysregulation (e.g., NMDA encephalitis) → Neuroinflammation → AMS.

Causes of Acute Encephalopathy

Category	Causes	Notes
Metabolic	-Hypoglycemia (<40 mg/dL). -Hyponatremia (Na+ <120 mEq/L). -Hypercapnia (CO2 >60 mmHg). - Uremia (BUN >100 mg/dL).	Hypoglycemia: Rapid onset, reversible with glucose.
Infectious	-Sepsis (e.g., E. coli bacteremia). -Meningitis (bacterial, viral). -Encephalitis (HSV, West Nile).	Sepsis: Most common in ICU; 70% have AMS.
Toxic	-Alcohol withdrawal (delirium tremens). -Opioid overdose (respiratory depression). -Medications: Benzodiazepines, anticholinergics.	Alcohol: Onset 48-72h post- cessation.
Structural	-Stroke (ischemic/hemorrhagic). -Intracranial hemorrhage (ICH). -Mass lesion (tumor, abscess).	Stroke: Focal deficits + AMS.

Category	Causes	Notes
Autoimmune	-NMDA receptor encephalitis. -Limbic encephalitis (anti-LGI1).	NMDA: Psychiatric symptoms, seizures, young patients.
Psychiatric	-Catatonia (e.g., schizophrenia). -Acute psychosis (e.g., brief psychotic disorder).	Catatonia: Stupor, rigidity; mimics AMS.
Other	-Hepatic encephalopathy (HE; ↑ ammonia)Hypoxic-ischemic (post-cardiac arrest) Seizures (non-convulsive status).	HE: Common in cirrhosis; asterixis.
Rare	-Wernicke's encephalopathy (thiamine deficiency) Posterior Reversible Encephalopathy Syndrome (PRES)CJD (rapidly progressive).	Wernicke's: Triad (confusion, ataxia, ophthalmoplegia).

Clinical Presentation

• Symptoms:

- Confusion, disorientation, agitation (delirium).
- Lethargy, stupor, coma (severe).
- Hallucinations (toxic, withdrawal, NMDA encephalitis).

• Exam:

- Mental Status: CAM-ICU (delirium), GCS (coma; <8 → Intubate).
- Neurologic: Asterixis (HE), focal deficits (stroke), nystagmus (Wernicke's), dyskinesias (NMDA).
- Psychiatric: Catatonia (rigidity, mutism), psychosis (delusions, disorganized behavior).
- **Systemic:** Fever (sepsis), jaundice (HE), respiratory depression (opioids).
- Substance Use: Alcohol withdrawal (tremors, seizures), opioid overdose (pinpoint pupils, bradypnea).
- Red Flags: GCS <8, unequal pupils (ICH), fever + neck stiffness (meningitis), dyskinesias + AMS (NMDA).

Diagnostic Workup

Initial Steps:

- Vitals: Hypoxia (Sp02 <90%), fever (>38°C), BP (PRES: >180/110 mmHg).
- Fingerstick Glucose: Hypoglycemia (<40 mg/dL) → D50W 50 mL IV.
- Labs:
 - Metabolic: CMP (Na+, BUN, glucose), ABG (CO2, pH), ammonia (HE).
 - Infectious: CBC, blood cultures, UA, CXR (sepsis).
 - **Toxic:** Urine drug screen (opioids, benzos), alcohol level, salicylate/acetaminophen levels.

- **Autoimmune:** CSF NMDA receptor antibodies, serum paraneoplastic panel (e.g., ovarian teratoma in NMDA).
- Other: TSH (hypothyroidism), B12, thiamine (Wernicke's).

Imaging:

- CT Head (Non-contrast): Stroke, ICH, mass lesion (1st step).
- MRI Brain: Encephalitis (HSV: temporal lobe), PRES (posterior edema),
 NMDA (normal or nonspecific).

Other Tests:

- Lumbar Puncture (LP): If meningitis/encephalitis suspected (after CT); CSF:
 Cell count, glucose, protein, Gram stain, HSV PCR, NMDA antibodies.
- EEG: Non-convulsive status epilepticus (NCSE; 20% of ICU AMS), NMDA (extreme delta brush pattern).
- **EKG:** Arrhythmia (hypoxic-ischemic).
- Psych Consult: If catatonia or psychosis suspected (e.g., Bush-Francis Catatonia Scale).
- **Key Tip:** Consider NMDA encephalitis in young patients with psychiatric symptoms + seizures; psych consult for catatonia mimics.

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Evaluation Flowsheet: Acute Encephalopathy

- Step 1: ABCs + Vitals: Hypoxia → O2; fever → Cultures; BP >180/110 → PRES.
- **Step 2:** Glucose: <40 mg/dL → D50W 50 mL IV.
- Step 3: History/Exam:
 - Focal deficits → CT (stroke).
 - Asterixis, jaundice → Ammonia (HE).
 - Tremors, seizures → Alcohol withdrawal.
 - Dyskinesias, hallucinations → NMDA encephalitis.
 - Rigidity, mutism → Psych consult (catatonia).
- Step 4: Labs/Imaging:
 - CMP, drug screen, cultures → Metabolic/toxic/infectious.
 - CT/MRI → Structural (stroke, ICH).
- Step 5: Advanced Testing:
 - LP → Meningitis (WBC >5/ μ L, \downarrow glucose) or NMDA (anti-NMDA antibodies).
 - EEG → NCSE or NMDA (delta brush).
 - Psych consult → Catatonia (lorazepam challenge).
- Step 6: Treat underlying cause: Sepsis → Antibiotics; HE → Lactulose; NMDA → IVIG; catatonia → Lorazepam.

Examples

- Case 1: Hepatic Encephalopathy (Cirrhosis)
- **Presentation:** 60 y/o M, cirrhosis, confusion, asterixis, ammonia 90 µmol/L (normal <30), no fever, CT normal.
- Interpretation: Acute encephalopathy (HE, ↑ ammonia), West Haven Grade II.
- Workup: Ammonia, CMP (bilirubin 4 mg/dL, INR 1.8), rule out infection (cultures, paracentesis).
- Next Steps: Lactulose (30 mL q2-4h until 2-3 BM/day), rifaximin (550 mg BID), identify trigger (e.g., GI bleed).
- Case 2: Septic Encephalopathy (UTI)
- **Presentation:** 75 y/o F, ICU, fever 39°C, lethargy, UA: +LE, +nitrites, blood cultures: E. coli.
- **Interpretation:** Acute encephalopathy (sepsis-related), CAM-ICU positive (delirium).
- Workup: CBC (WBC 15K), CMP (normal), CT head (normal), EEG (diffuse slowing).
- Next Steps: Antibiotics (ceftriaxone 2 g IV q24h), fluids (NS 30 mL/kg), monitor for NCSE.

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