Speaker: Allan Tunkel, MD





CASE #1

- 24-year-old female who presented with pain and swelling on the right side of her jaw that had been progressing over the last several weeks. She was unable to open her mouth. She denied fever or headache, and had no past hospitalizations or illnesses. The patient had not been to the dentist within 10 years.
- T 99.8°F, P 88, RR 14, BP 110/80
- Exam revealed swelling and erythema along her right mandible





Question #1 (Case #1)

Which of the following empiric antimicrobial regimens should be initiated?

- A. Ceftriaxone + metronidazole
- B. Vancomycin + cefepime
- c. Trimethoprim-sulfamethoxazole
- D. Voriconazole
- E. Liposomal amphotericin B

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PREDISPOSING CONDITIONS FOR BRAIN ABSCESS	
Condition	Relative Frequency (%)
Contiguous focus of infection (otitis media, mastoiditis, sinusitis, face or scalp infection, dental sepsis, osteomyelitis, penetrating head injury)	30-50
Hematogenous spread (lung abscess, empyema, congenital heart disease, bronchiectasis, infective endocarditis, compromised host, hereditary hemorrhagic telangiectasia)	~35
Cryptogenic	10-35

PRINCIPLES OF BRAIN ABSCESS MANAGEMENT

- MR imaging is the diagnostic procedure of choice; diffusion-weighted imaging increases diagnostic accuracy (sensitivity and specificity 96% for differentiation from cancers [PPV 98%; NPV 92%])
- Lumbar puncture is contraindicated
- Biopsy or aspiration (via stereotactic guidance) is needed for microbiologic diagnosis
- Begin empiric antimicrobial therapy based on underlying condition and pathogenesis of spread of infection to brain





EMPIRIC ANTIMICROBIAL THERAPY OF BRAIN ABSCESS

etronidazole + a third-generation cephalosporin ^a ancomycin + metronidazole + a third-generation aphalosporin ^a wird-generation cephalosporin ^a + metronidazole
ancomycin + metronidazole + a third-generation ephalosporin ^a nird-generation cephalosporin ^a + metronidazole
hird-generation cephalosporin ^a + metronidazole
c
ancomycin + a third or fourth generation ephalosporin
hird-generation cephalosporin ^a + metronidazole + imethoprim-sulfamethoxazole
ancomycin ^b
ai Pi

EMPIRIC ANTIMICROBIAL THERAPY OF BRAIN ABSCESS

Predisposing Condition	Antimicrobial Regimen
Unknown	Vancomycin + metronidazole + a third or fourth generation cephalosporin
Transplant recipients	Add voriconazole, plus trimethoprim- sulfamethoxazole or sulfadiazine
HIV-infected patients	Add pyrimethamine + sulfadiazine; consider isoniazid, rifampin, pyrazinamide, and ethambutol for possible tuberculosis

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CASE #2

- 21-year-old member of a motorcycle gang thrown from his bike, and suffered a depressed skull fracture
- In the OR, a large subdural hematoma was evacuated
- Discharged in 5 days
- Returned by mother 5 days later because of bizarre behavior
- No headache, afebrile





CASE #3

- 78-year-old male with multiple myeloma on chronic prednisone therapy; underwent aortic valve replacement with a bioprosthesis 5 years earlier; presented with new-onset seizures
- T 100.4° F, P 96, RR 18, BP 110/70 mmHg; Exam (-)
- CT scan revealed multiple ring-enhancing lesions
- TEE no vegetations and normal bioprosthesis
- Empirically placed on vancomycin + ampicillin + gentamicin
- Blood cultures negative





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Question #2 (Case #3)

Which of the following antimicrobial regimens should be initiated?

- A. Penicillin + metronidazole
- B. Trimethoprim-sulfamethoxazole
- c. Daptomycin
- D. Liposomal amphotericin B + 5-FC
- E. Voriconazole



CASE #4

- 24-year-old injection drug user who, while injecting intravenous drugs with his girlfriend, fell out of the second story window of his apartment. When he did not return for 48 hours, she found him unresponsive on the ground and called fire rescue
- T 103°F, P 150, RR 32, BP 110/76 mmHg
- On exam, he was comatose without evidence of head trauma
- WBC 13,000/mm³, profound metabolic acidosis



Question #3 (CASE #4)

The most likely etiologic agent of the patient's CNS lesions is which of the following?

- A. Staphylococcus aureus
- B. Pseudomonas aeruginosa
- c. Nocardia asteroides
- D. Candida albicans
- E. Rhizopus arrhizus



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CASE #5

- 11-year-old boy with chronic granulomatous disease on chronic TMP-SMX therapy noted the onset of a mild headache which lasted 10 minutes.
- Two weeks later at a routine physician visit, the patient had no complaints and denied recurrence of the headache
- On examination, the patient had normal vital signs and a normal neurologic examination
- The physician ordered an MR imaging of the head





CASE #6

- 80-year-old male with CLL on chronic prednisone therapy presented to the VA Hospital with sepsis and ARDS. Course complicated by VDRF and multiple nosocomial infections, including candidemia for which he received 4 weeks of IV liposomal amphotericin B. After completing the course of therapy, he developed altered mental status
- □ T 101⁰ F, P 100, RR 20, BP 120/76
- Neurologic exam left-sided hyperreflexia and Babinski



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PRINCIPLES OF BRAIN ABSCESS MANAGEMENT

- Optimal management usually requires a combined medical and surgical approach (aspirate if >2.5 cm)
- Fungal brain abscess often requires combined medical and surgical therapy
- Initiate corticosteroids with evidence of cerebral edema or mass effect causing increased ICP

ANTIMICROBIAL THERAPY OF BRAIN ABSCESS

Organism	Antimicrobial Therapy
Actinomyces sp.ª	Penicillin G
Bacteroides fragilis ^a	Metronidazole
Enterobacteriaceaeª	Third or fourth generation cephalosporin
Fusobacterium sp.ª	Metronidazole
Pseudomonas aeruginosa	Ceftazidime or cefepime or meropenem
Staphylococcus aureus	Nafcillin, oxacillin, or vancomycin
Strep. milleri; a other streptococcia	Penicillin G

ANTIMICROBIAL THERAPY OF BRAIN ABSCESS	
Organism	Antimicrobial Therapy
Nocardia asteroides	Trimethoprim-sulfamethoxazole or sulfadiazine; combination therapy for immunocompromised patients and those failing standard therapy
Mycobacterium tuberculosis	lsoniazid + rifampin + pyrazinamide <u>+</u> ethambutol

ANTIMICROBIAL THERAPY OF BRAIN ABSCESS

Organism	Antimicrobial Therapy
Aspergillus sp.	Voriconazole
Candida sp.	Lipid formulation of amphotericin B ^a
Mucorales	Lipid formulation of amphotericin B
Scedosporium spp.	Voriconazole

CASE #7	2022 PREVIEW QUESTION
 79-year-old female is trans failure to thrive as a result nasogastric tube is placed hyperalimentation 	ferred from a nursing home for of decreased oral intake. A via the left nares for enteral
 One week into her hospita fever to 101.5° F, and left 	l course, the patient develops periorbital edema and chemosis
 CT scan of the head without of the sphenoid sinus 	ut contrast reveals opacification

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Ques	tion #4 (CASE #7)
WI	hich of the following studies should be performed to establish the diagnosis?
А. В. С. D. Е.	CT scan of the head and sinuses with contrast MR imaging with MR venography Cerebral angiography Positron emission tomography of the head Lumbar puncture

EPIDEMIOLOGY AND ETIOLOGY OF SEPTIC CAVERNOUS SINUS THROMBOSIS

Risk Factors	Etiologic Agents
Paranasal sinusitis	Staphylococci (60-70%)
Facial infection	Streptococci (~17%)
Dental infection	Gram-negative bacilli (~5%)
	Pneumococci (~5%)
	Bacteroides sp. (~2%)

CLINICAL FEATURES OF SEPTIC CAVERNOUS SINUS THROMBOSIS

Symptoms	Signs
Headache (52%)	Periorbital edema (73%)
Facial pain	Chemosis
Vision loss	Papillitis
Fever	Oculomotor palsies
Double vision	Proptosis



RADIOLOGIC FINDINGS IN SEPTIC CAVERNOUS SINUS THROMBOSIS

MR imaging

- Noninvasive diagnostic procedure of choice
- MRA and MRV can directly visualize cerebral vasculature
- Fullness in cavernous sinus region
- Paranasal sinus fluid

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MANAGEMENT OF SEPTIC CAVERNOUS SINUS THROMBOSIS

- Culture and drainage of infected sinuses
- Antimicrobial therapy (vancomycin + metronidazole + 3rd or 4th generation cephalosporin)
- Anticoagulation
 - Cavernous sinus thrombosis
 - Lateral sinus thrombosis?
 - Superior sagittal sinus thrombosis?

CASE #8

- 22-year-old man with a history of paranasal sinusitis presents with fever, severe headache, neck pain, and seizure
- On physical examination, T 102° F and he is lethargic
- Laboratory studies normal



Question #5 (CASE #8)

In addition to appropriate antimicrobial therapy, what other management should be performed?

- A. Lumbar puncture
- B. External ventricular drain
- c. Dexamethasone
- D. Burr hole drainage
- E. Craniotomy

CRANIAL SUBDURAL EMPYEMA AND CRANIAL EPIDURAL ABSCESS

Risk Factors	Etiologic Agents
Sinusitis (50-80%)	Staphylococci (10-15%)
Otogenic	Streptococci (25-45%)
Head trauma	Gram-negative bacilli (3-10%)
Neurosurgery	Other anaerobes (8%)
Hematogenous	Others (8%)
Meningitis	Unknown (20%)

Epidural Abscess

(indolent course)

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CRANIAL SUBDURAL EMPYEMA AND CRANIAL EPIDURAL ABSCESS

Subdural Empyema

(acute course) (indo er Headache

- Fever
- Headache

Seizures

- Depressed consciousness
 - SeizuresFocal neurologic signs
 - Altered mental state

Fever

Nuchal rigidity

Hemiparesis

Gaze palsies/ataxia

PRINCIPLES OF MANAGEMENT OF CRANIAL SUBDURAL EMPYEMA

- MR imaging (diagnostic procedure of choice) provides better clarity of detail and can differentiate empyema from most sterile effusions and chronic hematomas; diffusionweighted imaging adds to value of MRI
- Surgical therapy (burr holes or craniotomy) is imperative; better outcome with craniotomy
- Empiric antimicrobial therapy based on pathogenesis of infection

SURGICAL MANAGEMENT OF CRANIAL SUBDURAL EMPYEMA		
Surgical Procedure	Mortality Rate	
Burr hole(s)	23.3%	
Craniectomy	11.5%	
Craniotomy	8.4%	
Nathoo et al. Neurosurgery 2001;49:872		



EPIDEMIOLOGY OF SPINAL EPIDURAL ABSCESS

- Usually occurs secondary to hematogenous dissemination (~50% of cases)
- $\hfill\square$ Contiguous foci (~1/3rd of cases)
- Unidentified source (20-40% of cases)
- Diabetes mellitus identified in up to 50% of patients

ETIOLOGY OF SPINAL EPIDURAL ABSCESS

Organism	Relative Frequency (%)
Staphylococci	50-90
Streptococci	8-17
Gram-negative bacilli	12-17
Other anaerobes	2
Other	2
> 1 organism	5-10
Unknown	6

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CLINICAL STAGES OF SPINAL EPIDURAL ABSCESS

- Back pain and tenderness at the level of infection
- Radicular pain and paresthesias
- Impaired spinal cord function; motor paresis and sensory deficits
- w. Complete paralysis

PRINCIPLES OF MANAGEMENT OF SPINAL EPIDURAL ABSCESS

- MR imaging is the diagnostic procedure of choice; can visualize the spinal cord and epidural space, and can identify accompanying osteomyelitis, intramedullary spinal cord lesions, and joint space infection
- Empiric antimicrobial therapy should include an antistaphylococcal agent and coverage for gramnegative bacilli

PRINCIPLES OF MANAGEMENT OF SPINAL EPIDURAL ABSCESS

- Surgical therapy imperative in the presence of neurologic dysfunction (best if <24-36 hours of complete paralysis)
- Nonsurgical therapy only for patients with an unacceptably high surgical risk or no neurologic deficits at diagnosis; patient must be followed carefully for clinical deterioration

QUESTIONS

Allan R. Tunkel, MD, PhD, MACP Email: <u>allan_tunkel@brown.edu</u>