

Infectious Diseases of the Central Nervous System

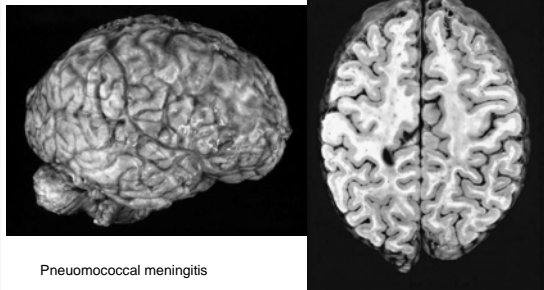
Bacterial Meningitis

Most common form of CNS infection

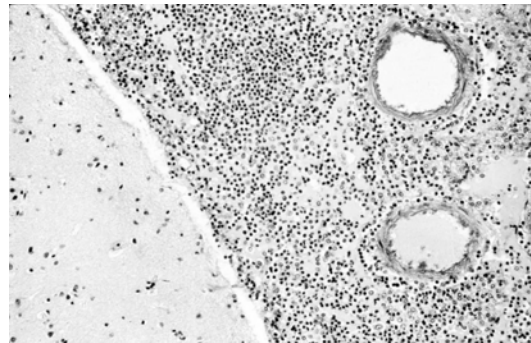
Organisms reach the leptomeninges via hematogenous spread or direct extension

Spinal tap yields cloudy CSF with many neutrophils and bacteria may be seen

Age group	Organism
Neonates	Group B streptococci; <i>E. coli</i>
Infants and children	<i>Haemophilus influenzae</i> (now <2 / 100,000)
Adolescents and young adults	<i>Neisseria meningitidis</i>
Elderly	<i>Streptococcus pneumoniae</i>



Pneumococcal meningitis



Brain abscess

Second most common infection of CNS following bacterial meningitis

Source of infection

Local contiguous spread (sinusitis, otitis, mastoiditis)

Hematogenous (Septic emboli from bacterial endocarditis, pulmonary infection, ect.)

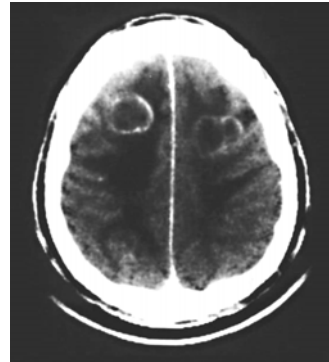
Stages of cerebral abscess formation

Early cerebritis (1-3 days)

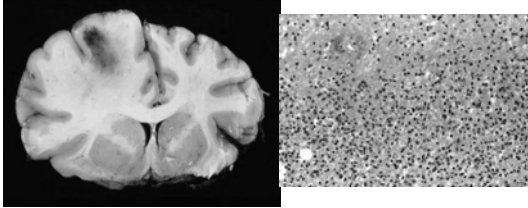
Late Cerebritis (4-9 days)

Early Capsule Formation (10-13 days)

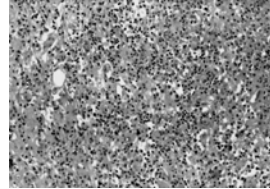
Late Capsule Formation (14 days and later)



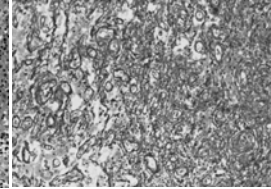
Early Cerebritis



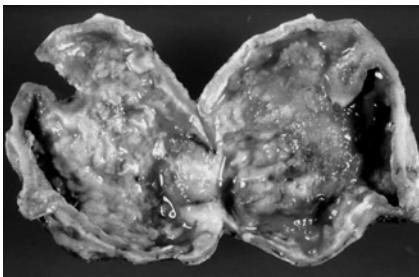
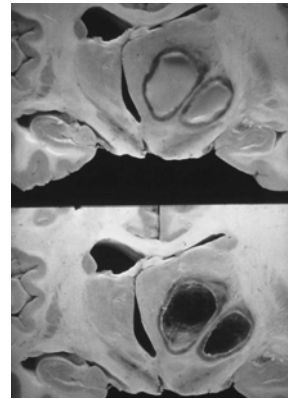
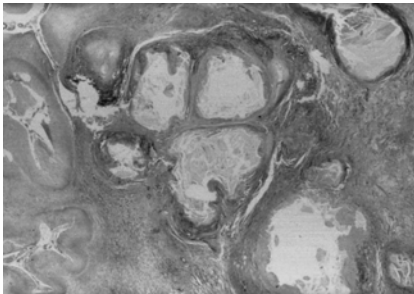
Late Cerebritis



Early Capsule Formation



Late Capsule Formation



Cerebral Fungal Infections

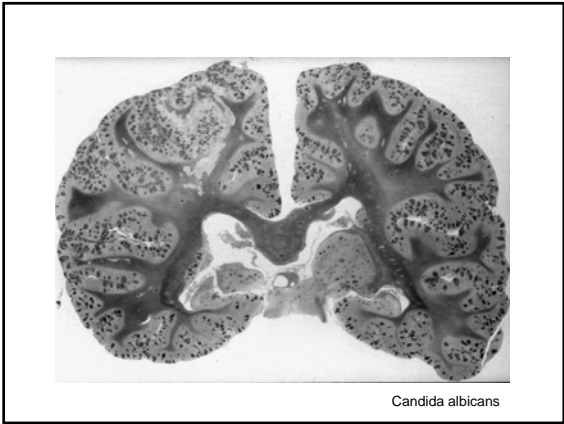
Often seen as an opportunistic infection in immunocompromised patients

Typically reach CNS via hematogenous spread from other organs

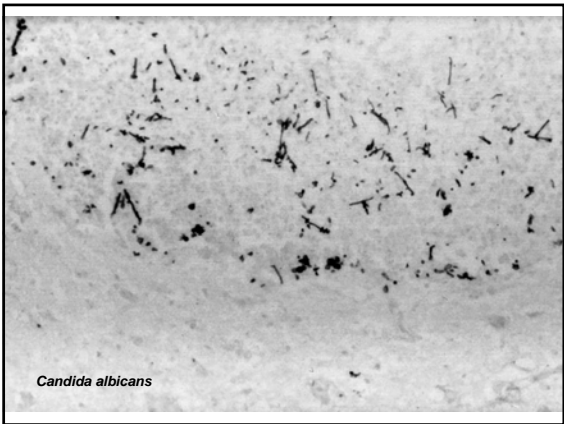
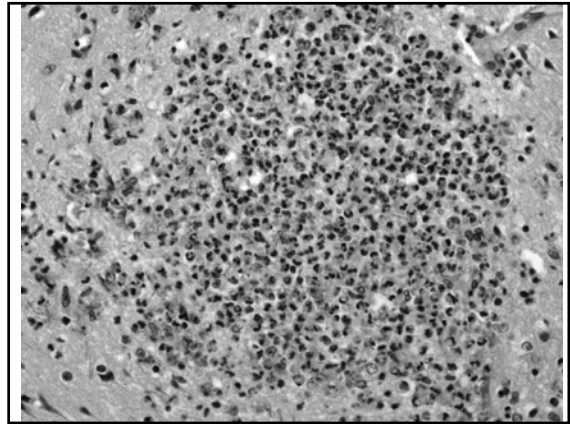
May produce leptomeningitis, vasculitis, granulomas or cerebral abscess

Common Organisms

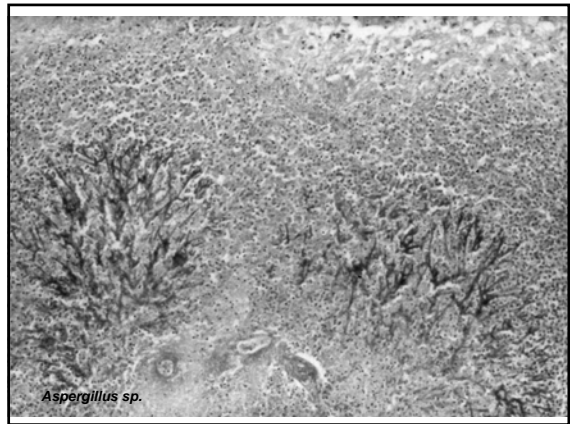
<u>Genus</u>	<u>Morphology</u>	<u>Patient status</u>
Aspergillus	Septate hyphae	Opportunistic
Mucormycosis	Nonseptate hyphae	Opportunistic
Candida	Budding yeast, pseudohyphae	Opportunistic
Cryptococcus	Budding yeast, encapsulated	Opportunistic or previously healthy



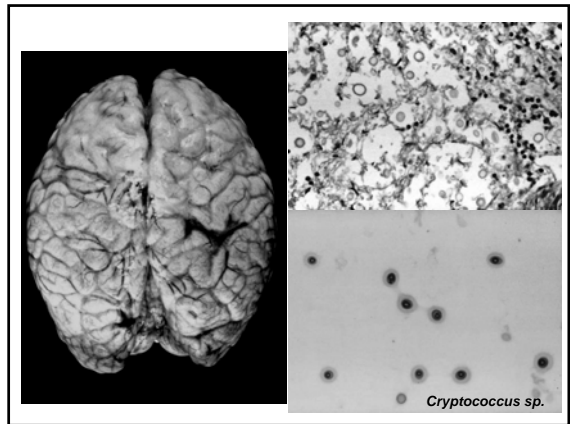
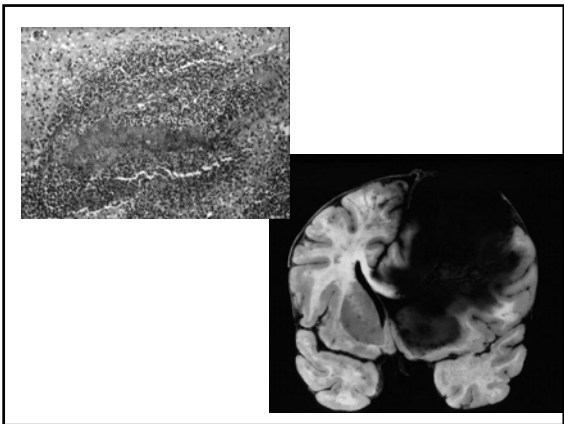
Candida albicans



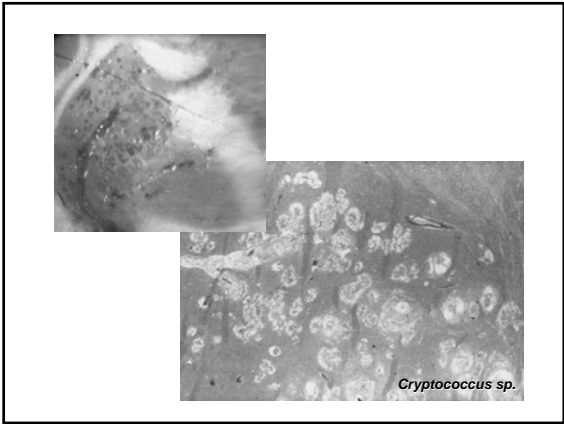
Candida albicans



Aspergillus sp.



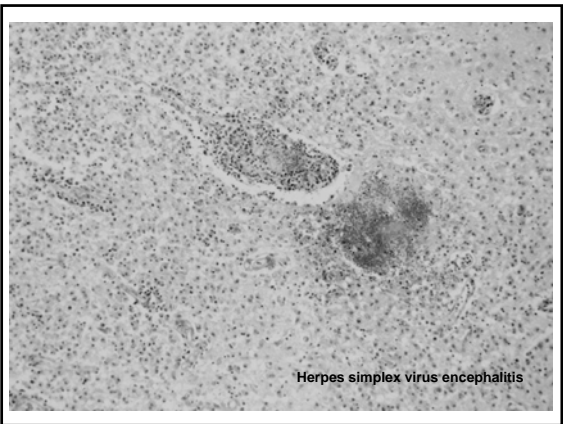
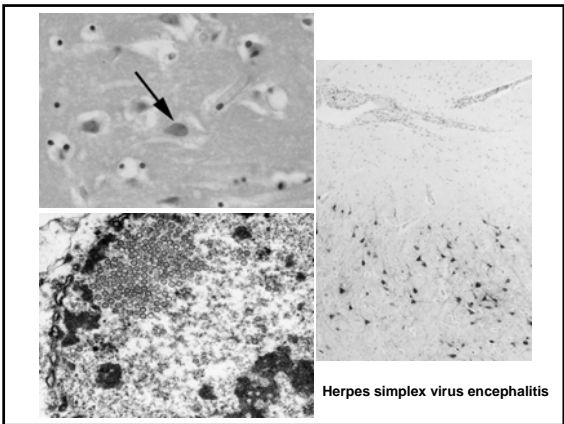
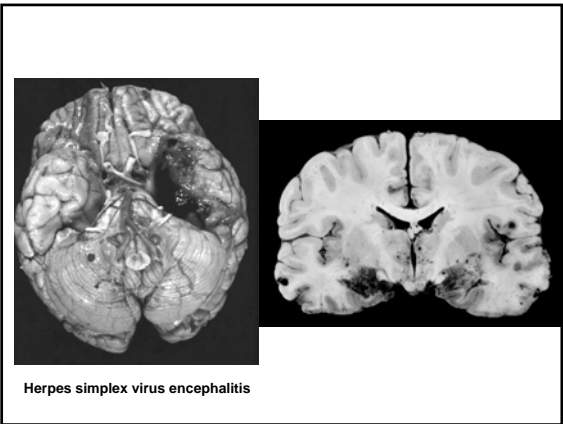
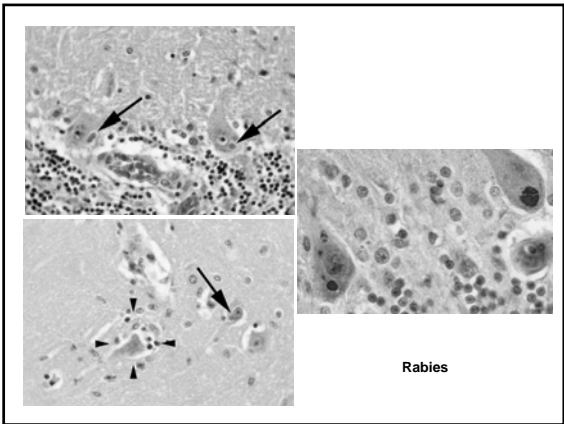
Cryptococcus sp.



CNS viral infections

Manifestations
 'Aseptic' meningitis
 Encephalitis
 Meningoencephalitis
 Myelitis

Stereotypical tissue reactions
 Inflammatory cell infiltrates
 Microgliosis
 Neuronophagia
 Microglial nodules
 Astrocytosis
 Intracellular inclusion bodies
 Neuronal cell degeneration
 Cellular and tissue necrosis



Neuropathology of AIDS

Human immunodeficiency virus type 1 (HIV-1)

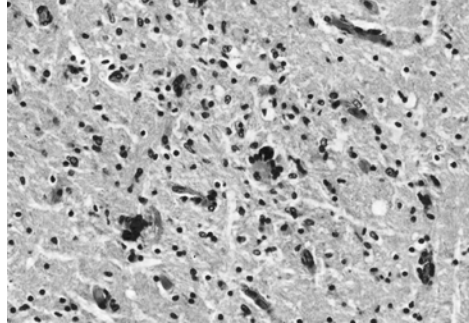
Primary complications

- HIV encephalitis or AIDS dementia complex
- HIV-associated myelopathy (vacuolar myelopathy)
- HIV-associated neuropathy (distal sensory neuropathy)
- HIV-associated myopathy

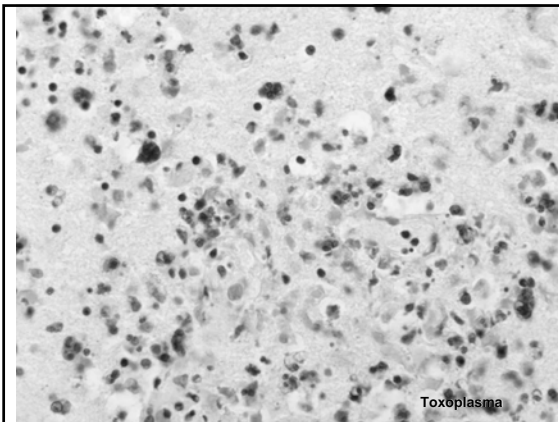
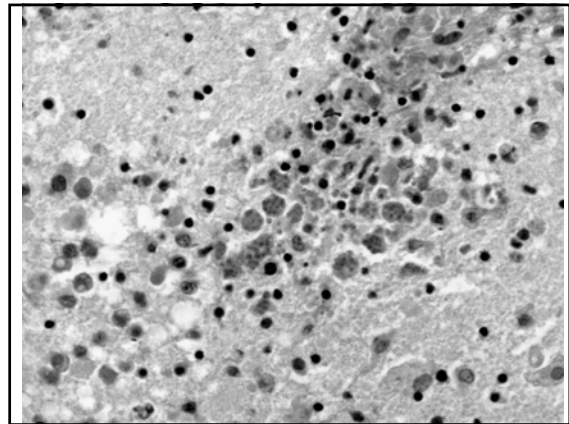
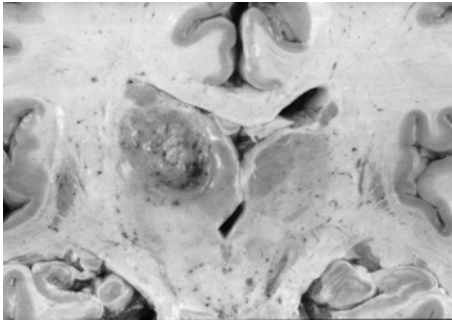
Secondary complications

- Opportunistic infections
 - Cryptococcosis
 - Toxoplasmosis
 - Progressive multifocal leukoencephalopathy
 - Cytomegalovirus infections
- Primary CNS lymphoma

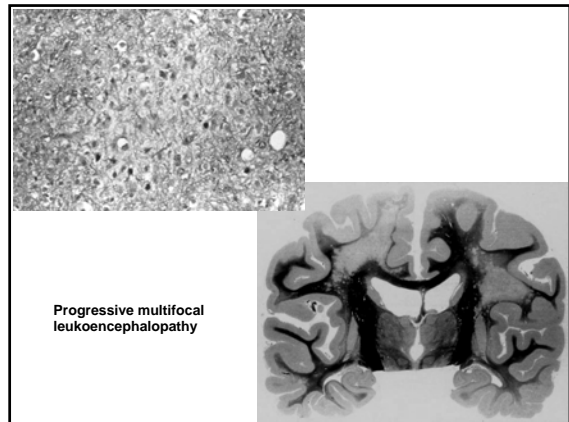
Microglial nodule with multinucleated giant cell



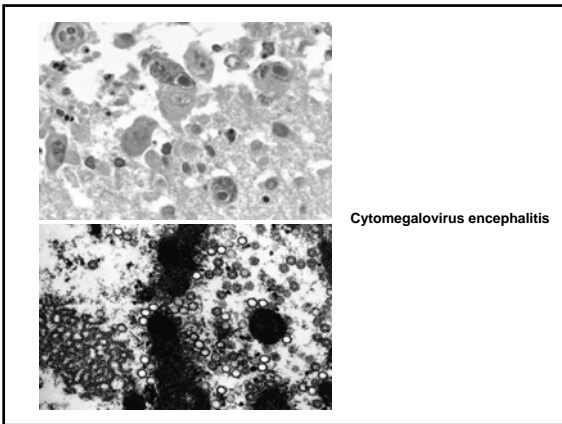
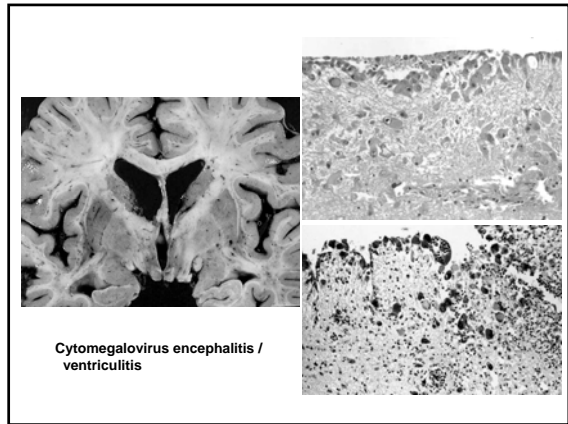
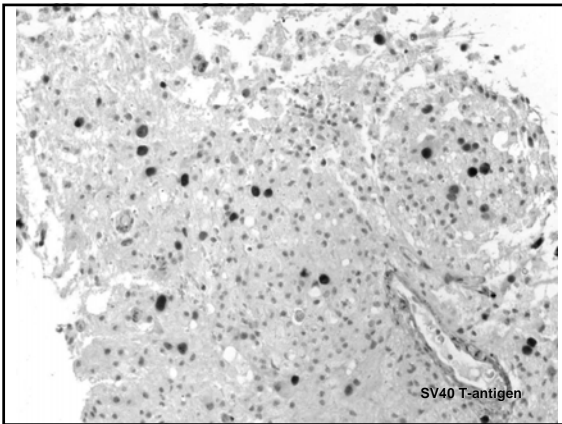
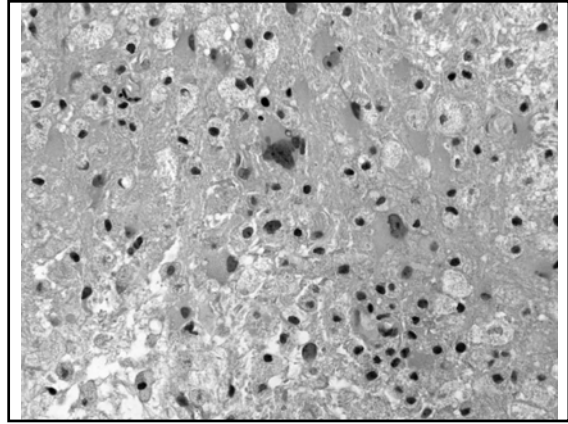
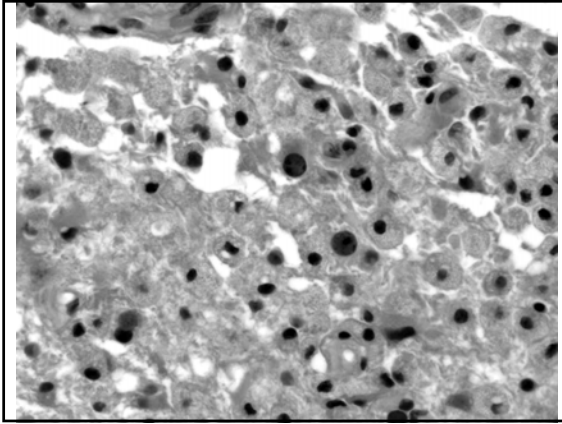
Large necrotic Toxoplasma lesion



Toxoplasma



Progressive multifocal leukoencephalopathy



Transmissible spongiform encephalopathies - Prion diseases

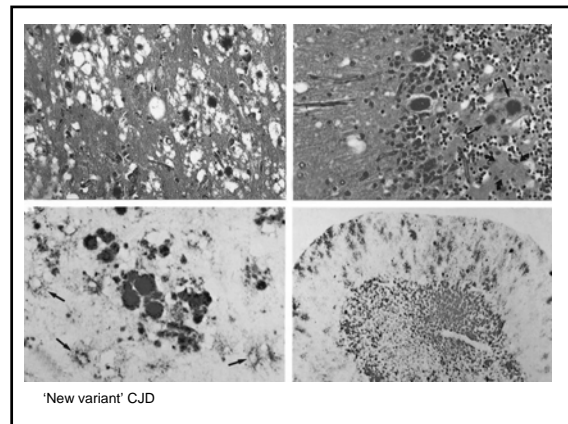
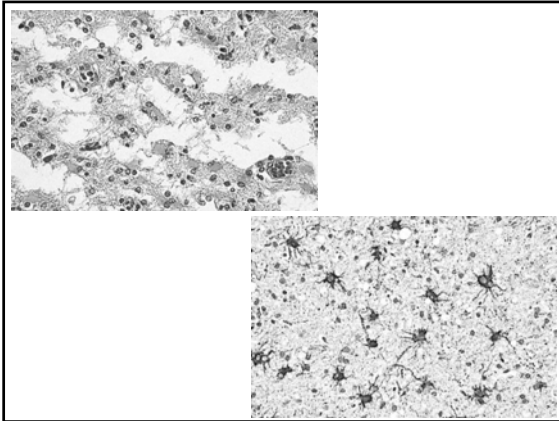
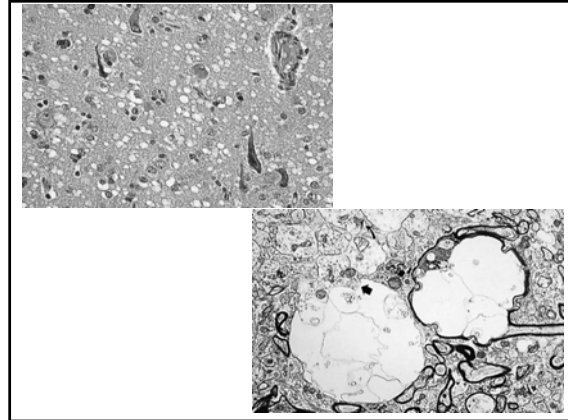
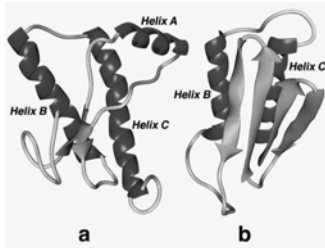
Creutzfeldt-Jakob Disease
 Worldwide incidence of approximately 1 per million
 Peak incidence in seventh decade of life
 Sporadic (85%), familial (15%) or iatrogenic transmission (very rare)
 Rapid progressive dementia, myoclonus, ataxia, usually fatal < 1 year

Other Human Prion Diseases
 Gerstmann-Straussler-Scheinker disease
 Fatal familial insomnia
 Kuru
 New Variant CJD (Mad Cows Disease)

Animal Prion Diseases
 Scrapie
 Bovine spongiform encephalopathy
 Others

Prion Hypothesis

PrP is a 30-KD normal cellular protein present in neurons
Disease occurs when PrP undergoes conformational change to a protease resistant form
This change occurs spontaneously at a very low rate - resulting in the sporadic cases
Various mutations in PrP facilitate the conformational change - familial cases
The infectious nature comes from ability of PrP^{Sc} to convert normal PrP
How accumulation of PrP^{Sc} causes neuronal cell death is still not understood



Other infections of the CNS

Arbovirus infections (arthropod-borne)
Poliomyelitis
Neurosyphilis
Neuroborreliosis (Lyme Disease)
Tuberculosis
Cysticercosis