• **Metastasis Cascade**
  - Invasion of *Basement Membrane* --> Invasion of *Extracellular Matrix* --> *Intravasation* (get into flowing lymph/blood) --> *Embolization* (float through vessels) --> *Extravasation* (lodge somewhere along course) --> *Angiogenesis/Growth*  

• **Dormancy** – period after metastasis lodges before growth occurs (“seeds” of metastasis aren’t growing yet)

• **Metastasis Appearance** – looks like cells from primary tissue, not tissue where metastasis lands!
  - **QUIZ:** Think about age/gender/etc of patient to get clues about where primary might be
  - Ex: *squamous cell carcinoma* of larynx – much more likely if middle-aged smoker
  - Look for either *squamous cell* tissue or *glandular “adenocarcinoma”* tissue

• **Metastasis Pathways**
  - **Lymphatic** – often causes *enlarged lymph nodes* (but enlargement not indicative of metastasis!)
  - **Hematogenous** – slides show tumor metastases from GI tract, testicle, breast
    - Pathologic Fracture – eating away at bone by metastasis, causing fracture = *osteolytic*
    - Osteoblastic Metastases – can cause bone growth
  - **Direct** – invade directly through tissue
    - Diaphragm underside from ovarian cancer