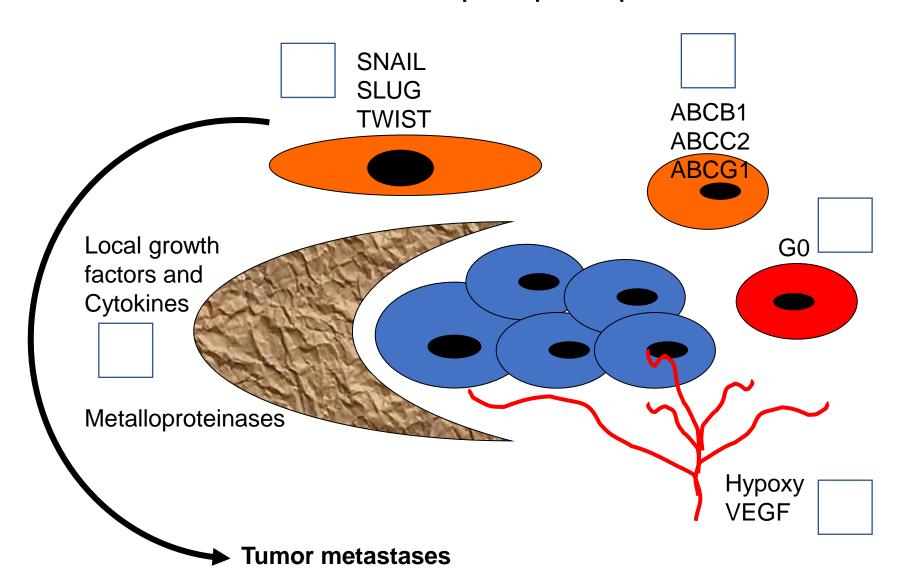
### Match the terms in a proper position...



#### Terms:

1. MDR Multidrug Resistance

2. Tumor stem cell

3. Tumor angiogenesis

**4. EMT**Epithelialmesenchymal
transition

5. Tumor stroma

#### Which metalloproteinases are most likely secreted by indicated cells?

- **Collagenases** MMP-1,-8,13,-18 are capable of degrading triple-helical fibrillar collagene of bone, cartilage, dentin in particular: I, II, III, V, IX
- **Gelatinases** , MMP-2 i -9 collagen type IV, laminin, gelatin
- Stromelysins, MMP-3 i -10 degradation of ECM
- Matrilysin MMP-7 i -26 degradation of matrix + celavage of: FASL, pro TNFalfa, E-cadherin
- Membrane type MMP are localized directly in the cell membrane
- Other MMP
- TIMP tissue inhibitos of metalloproteinases

TIMP 1, 2, 3, 4

2. Membane bound MMPs **Macrophages Neutrophils** 3. TIMPs **Pericytes** 4. All types of MMPs Intratumoral fibroblasts Cancer cells undergoing EMT **Fibroblasts** at the invasive front

1. Gelatinases

# Epithelial-mesenchymal transition (EMT) – discuss the phenomenon

- Cells of epithelial phenotype with strong cell-cell connections (kadherins), and strong cell ECM connections (integrins) gain the ability to move
- These cells undergoing EMT loose their connections with ECM.
- EMT is accompanied with change in cell morphology form epithelial like to mesenchyme like
- Changes in kadherin expression occurs (N-cadherin is up-regulated while E-cadhetin is repressed)

# Tumor angiogenesis – match the term with its definition

Vasculogenesis:

 de novo formation form angioblasts

Angiogenesis:

 basing on existing blood vessels, by proliferation of endothelium

• Vasculogenic mimicry:

 is the formation of microvascular channels by aggressive, metastatic and genetically deregulated tumur cells

 a small molecular inhibitor of several tyrosine protein kinases that inhibit mainly tumor angiogenesis

• This drug is approved for the treatment of primary kidney cancer (advanced renal cell carcinoma), advanced primary liver cancer (hepatocellular carcinoma), and radioactive iodine resistant advanced thyroid carcinoma.

## Targeted drugs – fill in the gaps

**Imatynib** 

• Tyrosine kinase inhbitiors .....

**Everolimus** 

• B-Raf inhibitors – .....

Bevacizumab

• mTOR inhbibirots – ....., tacrolimus (immunosupresive tacrolimus)

**Trastuzumab** 

• Pan kinase inhibitor angiogenesis inhibitor ......

Vemurafenib

• HER2 inhibitor –.....

Sorafenib

• Proteasome inhbibitor – .....

• Estrogene receptor inhibitor — ......

Bortezomib, carfilzomib

• **VEGF-A** ab — .....(avastin)

## Anticancer drugs

Cell-cycle unspecific

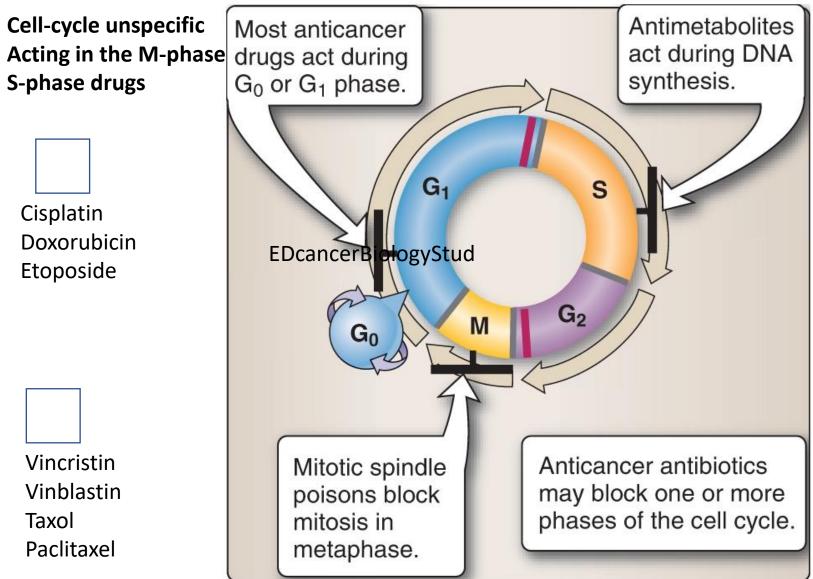
S-phase drugs



Cisplatin Doxorubicin Etoposide



Vincristin Vinblastin Taxol **Paclitaxel** 





Fluorouracyl Hydroxyurea Methotrexate