### Summary of cell grammar for PhysiCell (signals and behaviors)

# Key words:

- cell type: refers to a specific cell type (cell definition) previously defined in PhysiCell
- substrate: refers to a specific substrate previously defined in Physicell

#### Signals:

- 0 : substrate
- 1: intracellular substrate
- 2 : substrate gradient
- 3: pressure
- 4: volume
- 5 : contact with cell type
- 6: contact with live cell
- 7: contact with dead cell
- 8 : contact with apoptotic cell
- 9: contact with necrotic cell
- 10 : contact with other dead cell
- 11 : contact with basement membrane
- 12: damage
- 13 : damage delivered
- 14: attacking
- 15: dead
- 16: total attack time
- 17 : time
- 18: custom:sample
- 19: apoptotic
- 20: necrotic

#### **Behaviors**

- 0: substrate secretion
- 1 : substrate secretion target
- 2 : substrate uptake
- 3 : substrate export
- 4 : cycle entry
- 5: exit from cycle phase 1
- 6: exit from cycle phase 2
- 7: exit from cycle phase 3
- 8: exit from cycle phase 4
- 9: exit from cycle phase 5
- 10: apoptosis
- 11: necrosis

- 12: migration speed
- 13: migration bias
- 14: migration persistence time
- 15 : chemotactic response to substrate
- 16: cell-cell adhesion
- 17 : cell-cell adhesion elastic constant
- 18 : adhesive affinity to cell type
- 19 : relative maximum adhesion distance
- 20 : cell-cell repulsion
- 21: cell-BM adhesion
- 22 : cell-BM repulsion
- 23 : phagocytose apoptotic cell
- 24 : phagocytose necrotic cell
- 25 : phagocytose other dead cell
- 26: phagocytose cell type
- 27: attack cell type
- 28: fuse to cell type
- 29: transition to cell type
- 30: asymmetric division to cell type
- 31: custom:sample
- 32: is\_movable
- 33: immunogenicity to cell type
- 34 : cell attachment rate
- 35 : cell detachment rate
- 36: maximum number of cell attachments
- 37 : attack damage rate
- 38: attack duration
- 39: damage rate
- 40 : damage repair rate

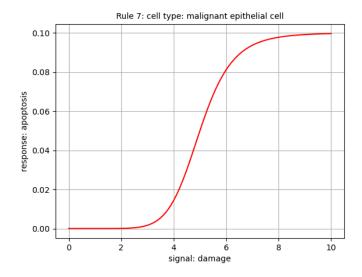
## Fundamental structure of the cell grammar:

Cell type | Signal | increases/decreases (also called Direction) | Behavior | Saturation Value | Half-max | Hill power | Apply to dead (Boolean Flag)

# **Examples:**

### 1) standard rule implementation

malignant epithelial cell | damage | increases | apoptosis | 0.1 | 5 | 8 | 0

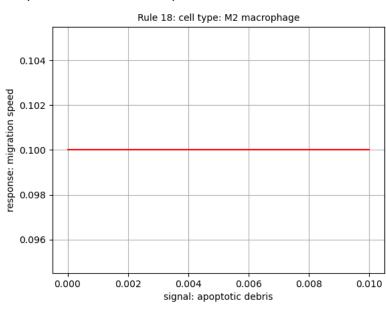


#### 2) English style rule implementation

In M2 macrophage cells:

apoptotic debris decreases migration speed from 1 towards 0.1 with a Hill response, with half-max 0.005 and Hill power 4.

... (more rules below)...



#### Tumor immune example rule set:

- 1. tumor,pressure,decreases,cycle entry,0,0.25,3,0
- 2. tumor, oxygen, increases, cycle entry, 0.0072, 21.5, 4,0
- 3. tumor, oxygen, decreases, necrosis, 0, 3.75, 8, 0
- 4. tumor,damage,increases,apoptosis,0.023,45,16,0
- 5. tumor, dead, increases, debris secretion, 0.017, 0.1, 10, 1
- 6. M0 macrophage, contact with dead cell, decreases, migration speed, 0.1, 0.1, 4, 0
- 7. M0 macrophage,contact with dead cell,increases,transform to M1 macrophage,0.05,0.1,10,0
- 8. M0 macrophage, dead, increases, debris secretion, 0.017, 0.1, 10, 1
- 9. M1 macrophage, contact with dead cell, decreases, migration speed, 0.1, 0.1, 4, 0
- 10. M1 macrophage, oxygen, decreases, transform to M2 macrophage, 0, 5, 4, 0
- 11. M1 macrophage, dead, increases, debris secretion, 0.017, 0.1, 10, 1
- 12. M2 macrophage, contact with dead cell, decreases, migration speed, 0.1, 0.1, 4, 0
- 13. M2 macrophage, dead, increases, debris secretion, 0.017, 0.1, 10, 1
- 14. naive T cell, IL-10, decreases, transform to CD8 T cell, 0, 0.25, 2, 0
- 15. naive T cell, IFN-gamma, increases, transform to CD8 T cell, 0.01, 0.25, 2, 0
- 16. naive T cell, dead, increases, debris secretion, 0.017, 0.1, 10, 1
- 17. CD8 T cell,IL-10,decreases,attack tumor,0,0.25,2,0
- 18. CD8 T cell,IL-10,decreases,migration speed,0.1,0.25,2,0
- 19. CD8 T cell, contact with tumor, decreases, migration speed, 0.1, 0.1, 2, 0
- 20. CD8 T cell, IFN-gamma, increases, cycle entry, 0.00041, 0.25, 2, 0
- 21. CD8 T cell,IL-10,increases,transform to exhausted T cell,0.005,0.25,4,0
- 22. CD8 T cell, dead, increases, debris secretion, 0.017, 0.1, 10, 1
- 23. exhausted T cell, dead, increases, debris secretion, 0.017, 0.1, 10, 1

#### Cell types:

- 1. tumor
- 2. M0 macrophage
- 3. M1 macrophage
- 4. M2 macrophage
- 5. naive T cell
- 6. CD8 T cell
- 7. exhausted T cell

#### Substrates:

- 1. oxygen
- 2. debris
- 3. IFN-gamma
- 4. IL-10