

Table Host Handout

In Vitro Lecture and Luncheon for Students



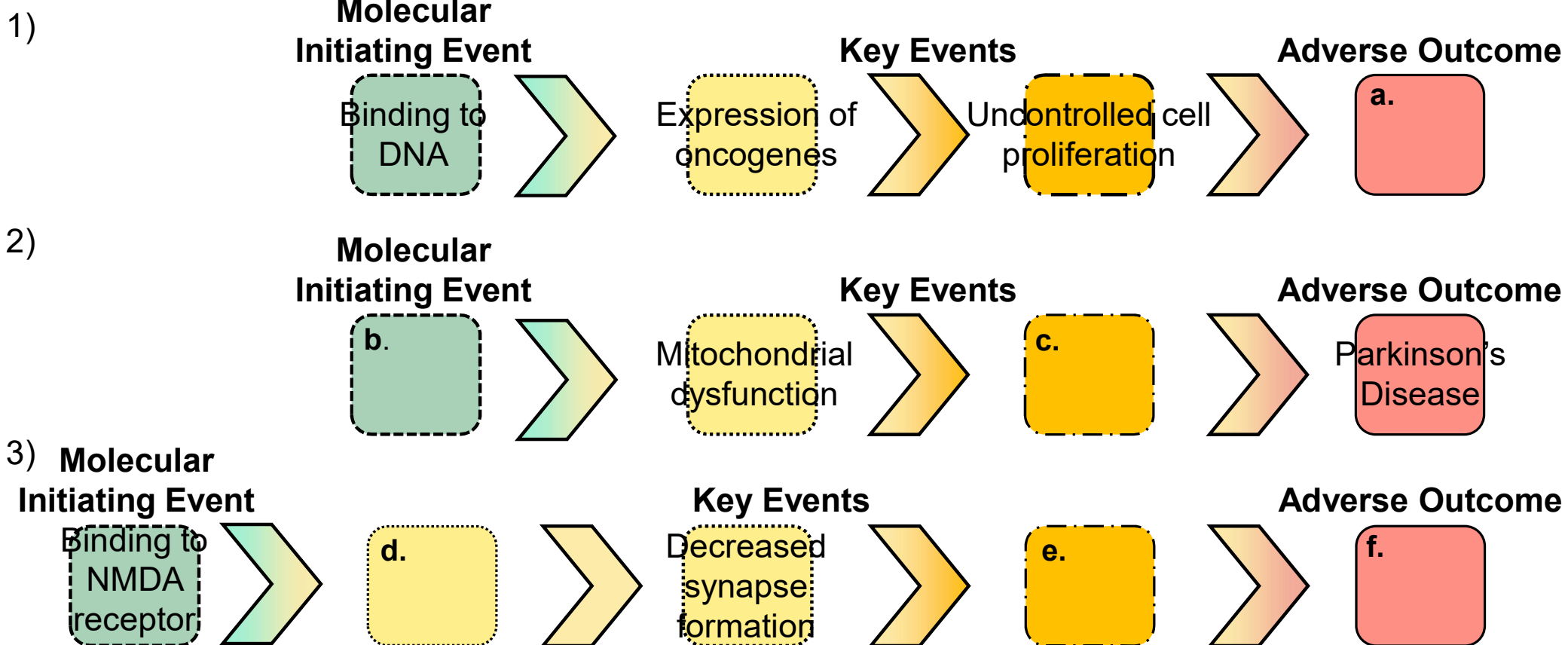
Making New Approach Methods Work
for Regulation: Linking Validation and
Biological Relevance Through Adverse
Outcome Pathways

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TABLE HOST—Exercise 1

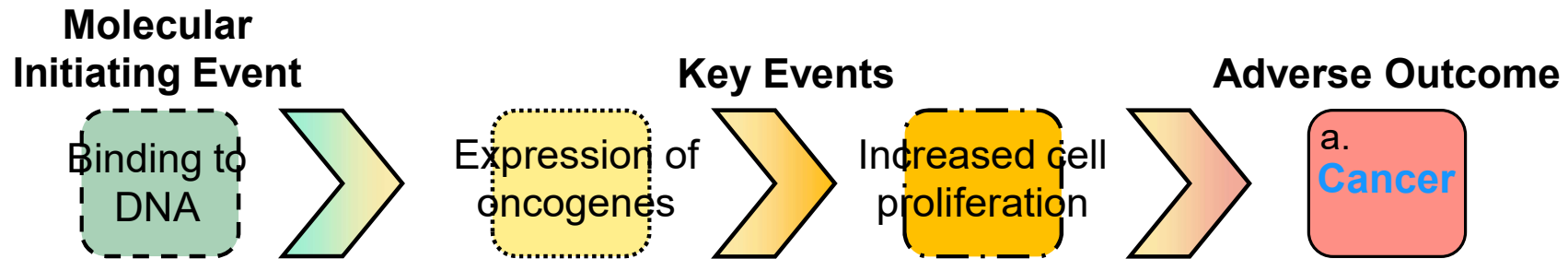
Match one of the six items on the right to each gap (a-f) in the three AOPs.

| | | |
|-----------------------------|---------------------------------------|------------------------|
| Apoptosis/ cell death | Binding to enzyme in the mitochondria | Cancer |
| Decreased neuronal function | Impaired learning and memory | Reduced calcium influx |



Exercise 1: Answers

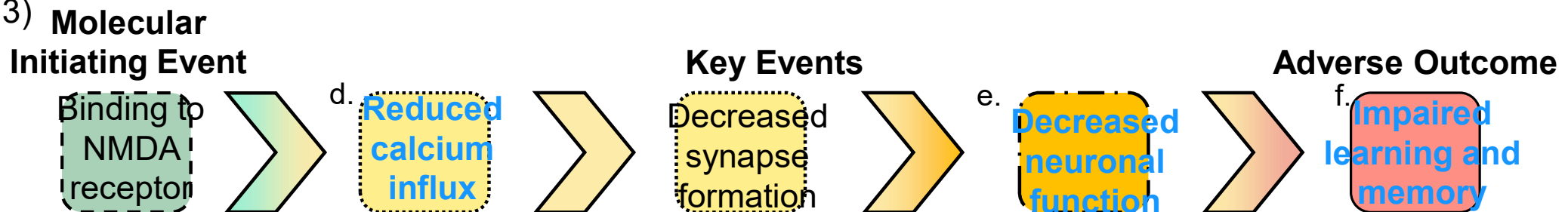
1)



2)



3)



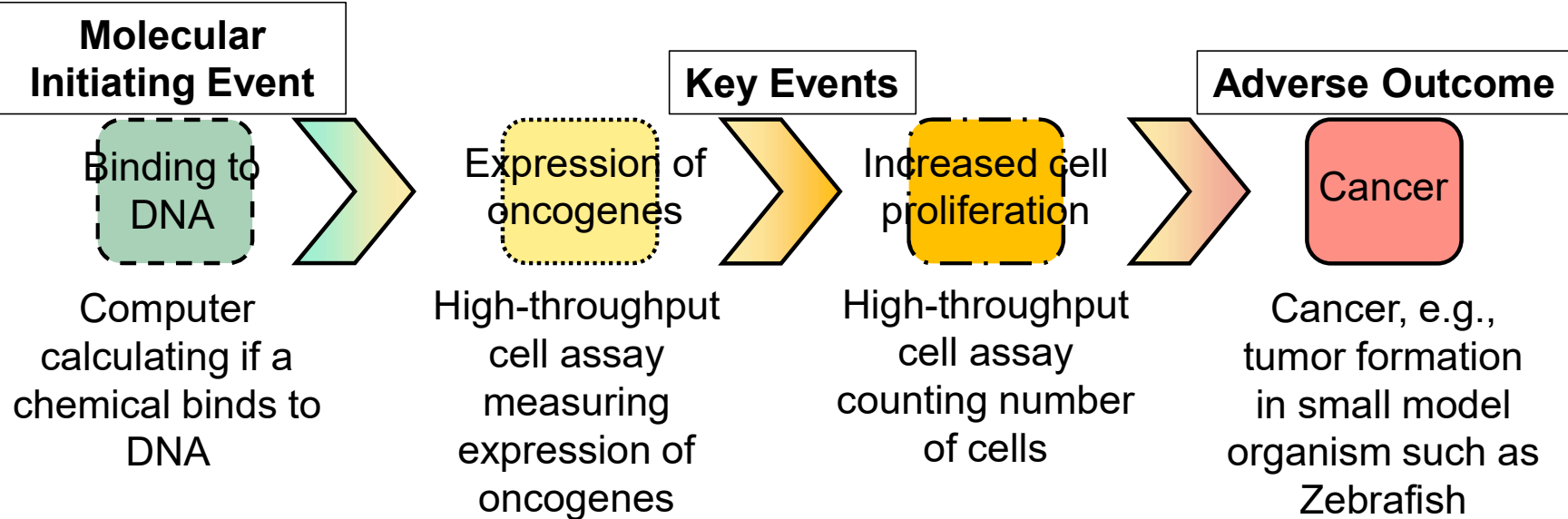
Exercise 2

Select one of the three AOPs in exercise 1, discuss and suggest types of NAMs (computational approaches, high throughput cell assays, 3D cell models, organs on chips, small model organisms), and what you could measure for the different events.



Exercise 2: Answers

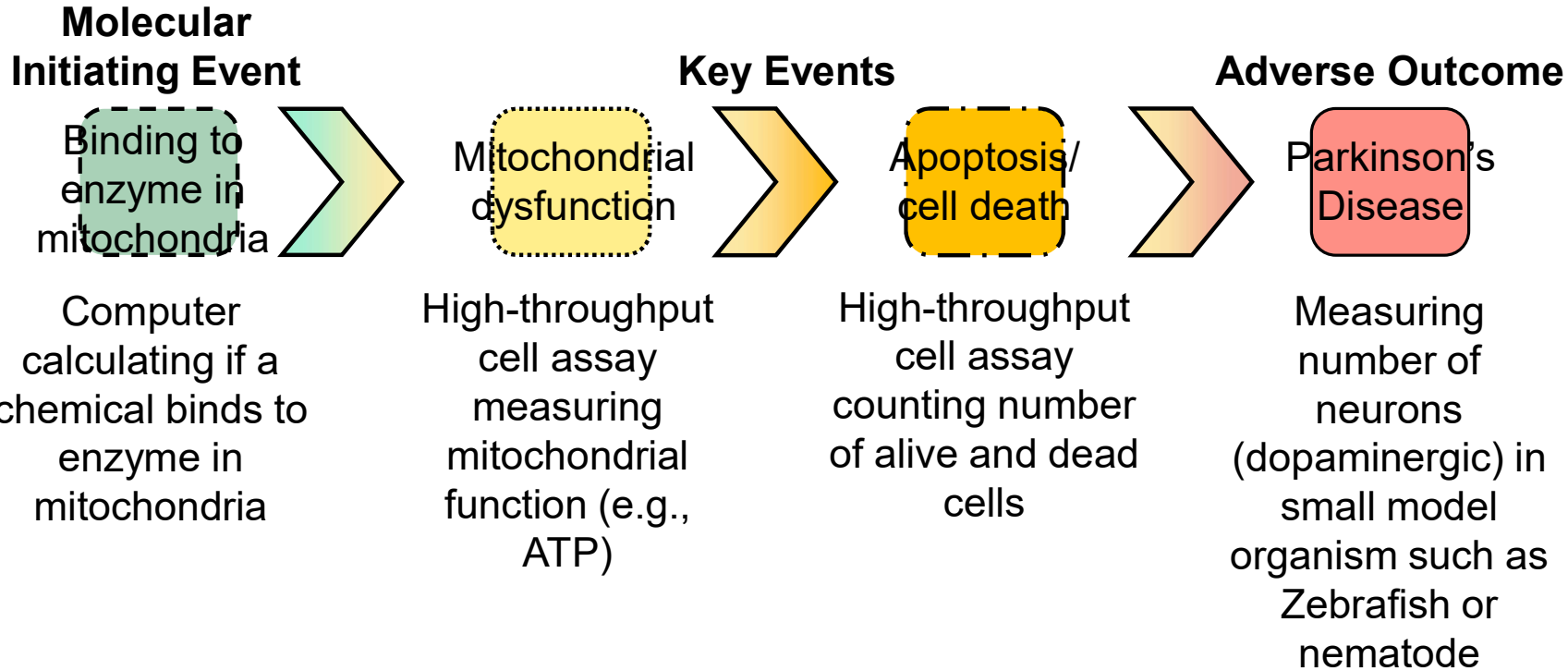
1)



Example NAM

Exercise 2: Answers

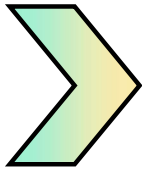
2)



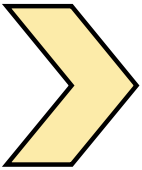
Exercise 2: Answers

3) Molecular Initiating Event

Binding to NMDA receptor

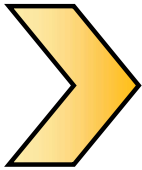


Reduced calcium influx

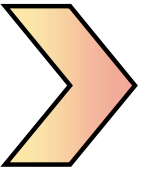


Key Events

Decreased synapse formation



Decreased neuronal function



Adverse Outcome

Impaired learning and memory

Computer model calculates if a chemical will bind to the NMDA receptor

High-throughput cell assay measures amount of calcium (e.g., by fluorescent dye)

High-throughput cell assay counting number of synapses

3D cell model on chip recording electrical activity of neurons (function)

Recording swimming behavior (distance and speed) of Zebrafish embryos