

Supplemental Video Legends

Supplemental Video 1: The accompanying animation to [Fig. 3](#).

The binding of (green) $150,000 \times 19 \times 10^{-9}$ m receptors to (red) $150,000 \times 21 \times 10^{-9}$ m ligands was simulated in 0.1 s increments under different conditions. Cells were represented by spheres with 5×10^{-6} m radii and the flat contact area had a radius of 4×10^{-6} m. The on-rate was $2.0 \times 10^5 \text{ M}^{-1} \text{ s}^{-1}$, the off-rate was 0.1 s^{-1} , and the confinement distance was 2×10^{-9} m. **(TOP LEFT)** Binding to an immobile receptor was simulated. Membranes were spaced 40×10^{-9} m apart. The receptors and their complexes with ligands were immobile while free ligands had diffusion coefficients of $3.0 \times 10^{-14} \text{ m}^2 \text{ s}^{-1}$. **(TOP MIDDLE)** Binding to a mobile receptor was simulated. The receptors now also had diffusion coefficients of $3.0 \times 10^{-14} \text{ m}^2 \text{ s}^{-1}$ while complexes had diffusion coefficients of $1.5 \times 10^{-14} \text{ m}^2 \text{ s}^{-1}$. **(TOP RIGHT)** Binding-induced receptor arrest was simulated. Free receptors and ligands had diffusion coefficients of $3.0 \times 10^{-14} \text{ m}^2 \text{ s}^{-1}$ while complexes had diffusion coefficients of $0.5 \times 10^{-15} \text{ m}^2 \text{ s}^{-1}$. **(BOTTOM LEFT)** The effects of suboptimal membrane separation were simulated. 100,000 virtual molecules led to a 20×10^{-9} instead of 40×10^{-9} m starting membrane separation. The σ for the normal curve penalizing the on-rate was 10×10^{-9} m. The σ for the normal curve penalizing movement between regions was 10×10^{-9} m for the 40×10^{-9} m complexes and 5.25×10^{-9} m for the 21×10^{-9} m ligands. **(BOTTOM MIDDLE)** The effects of a decreased σ on the movement of complexes were simulated. The σ for the normal curve penalizing the movement of complexes between regions was reduced to 5×10^{-9} m. **(BOTTOM RIGHT)** The effects of increasing the contribution of receptor/ligand complexes to the membrane separation were simulated. The numbers of virtual molecules were reduced to 20,000.

Supplemental Video 2: The accompanying animation to [Fig. 4](#).

150,000 molecules that moved towards the contact area with a velocity of $0.05 \times 10^{-6} \text{ m s}^{-1}$ and had diffusion coefficients of $1.0 \times 10^{-15} \text{ m}^2 \text{ s}^{-1}$ were simulated for 600 s in 0.1 s increments. Cells were represented by spheres with 5×10^{-6} m radii and contact areas were either curved or flat with radii of 4×10^{-6} m. The curved contact area was divided into 10 rings of 20 sectors. The flat contact areas were divided into 20×20 grids of square regions. Each region was allowed to contain up to 6250 molecules. The locations of molecules are shown when there was directed movement towards a **(LEFT)** curved or **(MIDDLE)** flat contact area where the targets were centered at the middle of the contact area and normally distributed with a σ of 0.1×10^{-6} m. **(RIGHT)** The locations of molecules are shown when there was directed movement towards a flat contact area where the targets were centered 3.5×10^{-6} m away from the middle of the 4×10^{-6} m contact area and were normally distributed with a σ of 0.5×10^{-6} m.

Supplemental Video 3: The accompanying animation to [Fig. 5](#).

The same conditions used in [Fig. 5](#) were used. The locations of **(TOP RED)** MHC complexes with off-rates of 0.02 s^{-1} , **(TOP GREEN)** MHC complexes with off-rates of 0.2 s^{-1} , **(TOP BLUE)** MHC complexes with off-rates of 2.0 s^{-1} , **(LEFT RED)** ICAM-1, **(LEFT BLUE)** CD48, **(RIGHT RED)** LFA-1, **(RIGHT GREEN)** TCRs and **(RIGHT BLUE)** CD2 are shown.