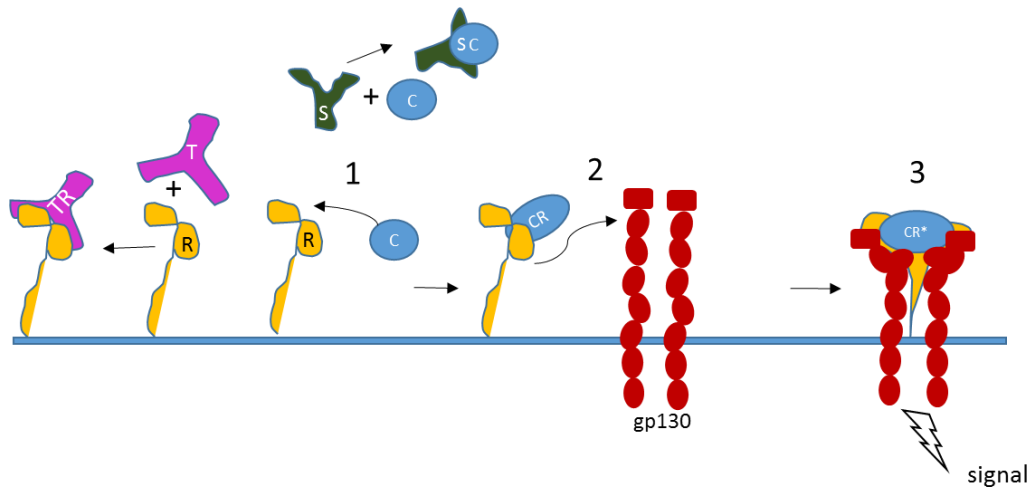


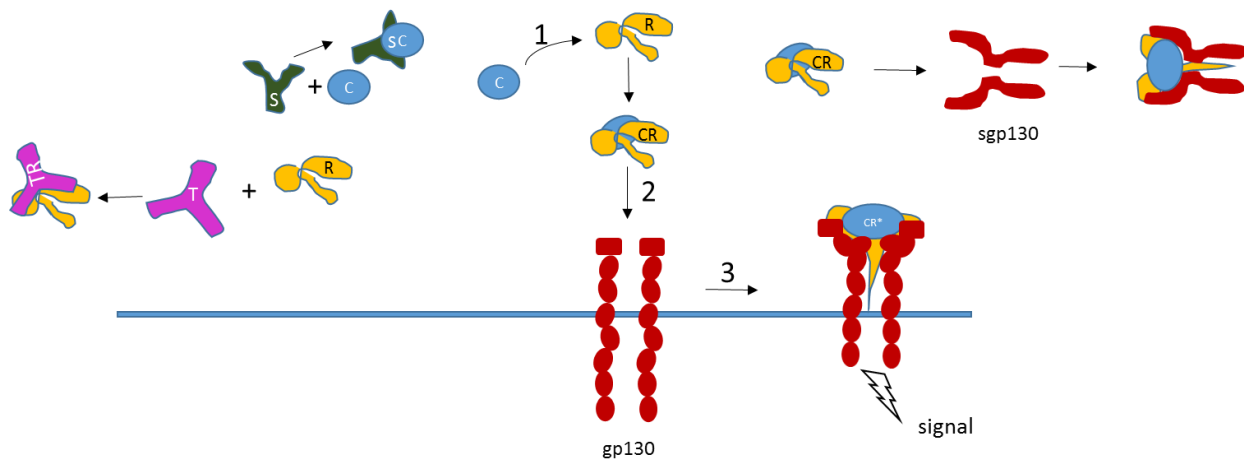
Figure 1: IL-6 Classic and Trans Signaling Pathways

(a) Classic Pathway



C: IL-6; **S:** siltuximab; **SC:** IL-6:siltuximab complex; **R:** IL-6R; **T:** TCZ; **TR:** IL-6R:TCZ complex; **CR:** IL-6:IL-6R pre-signaling complex; **CR*:** IL-6:IL-6R complex bound to gp130 (signaling complex)

(b) Trans Pathway



C: IL-6; **S:** siltuximab; **SC:** IL-6:siltuximab complex; **R:** sIL-6R; **T:** tocilizumab; **TR:** sIL-6R:TCZ complex; **CR:** IL-6:sIL-6R pre-signaling complex; **CR*:** IL-6:sIL-6R complex bound to gp130 (signaling complex)