TLR4 is activated mainly through the myeloid differentiation factor 88 (MyD88) signaling pathway. Signals from MyD88 are transmitted to the interleukin–1 receptor–associated kinase (IRAK) family of protein kinases via the interaction between MyD88 and the IRAK4 death domain [45] to recruit the tumor necrosis factor receptor–associated factor 6 (TRAF–6)[46–48]. Before ligand–induced signaling occurs, TLR4 first needs to associate with its extracellular binding partner, myeloid differentiation factor 2 (MD–2). [41, 42]. TLR4–MD2 dimerization recruits TIR domain–containing adaptor protein (TIRAP) and MyD88