Decrease heart rate and cardiac output How?– Acetylcholine binds to the M2 receptor in the heart activating Gi (GPCR), which will eventually lead to the hyperpolarization of the cell, because it opens the K+ channels and allows the K+ to leave the cell (makes the intercellular environment more negative).Recall that the Vagus nerve originates from cranial nerve X, which branches to the heart and ???.(lungs and the GI-tract (see figure