From Springfield:
I-55 South to IL Rt. 16 West
Rt. 16 West to Rt. 111 South
Rt. 111 South to Lewis and Clark

From Lambert International Airport:
I-70 East to I-170 North
North on I-170 to I-270 East
I-270 East to MO Rt. 367 North
MO Rt. 367 North to Rt. 67 North
Follow Rt 67/111 to Lewis and Clark

From Downtown St. Louis:
I-70 East to I-255 North
I-255 North to IL 143 North
IL143 North to IL 111 North
(143 turns into 67/111 at Clark Bridge in Alton)
Follow 111/67 North to Lewis and Clark

OR

From I-70, take Salisbury Street Exit (248A)
Cross McKinley Bridge, Follow Rt. 3 North to Berm Highway (Rt. 143)
Rt. 143 turns into 67 at Clark Bridge in Alton
Follow 67 North to Lewis and Clark

From Mt. Vernon:
I-64 West to I-255 North
I-255 North toWood River Exit (Rt. 143)
Turn Left (W) on 143 and continue
143 will turn into 67 at the Clark Bridge
Follow 67 North to Lewis and Clark