The majority of counties with high opioid-involved mortality rates are clustered in and to the south of the St. Louis metropolitan region, though some additional counties with high opioid-involved death rates can be found across the state. Missouri’s metropolitan areas have consistently high rates of opioid-involved overdose mortality. It should be noted that many counties’ death rates are based on fewer than 20 deaths and should be interpreted cautiously as mortality rates based on low counts can be unstable. Opioid-involved emergency room discharge rates follow a similar statewide distribution with the greatest rates in the St. Louis area; however, there is an additional cluster of high rates in the southwest region of the state.