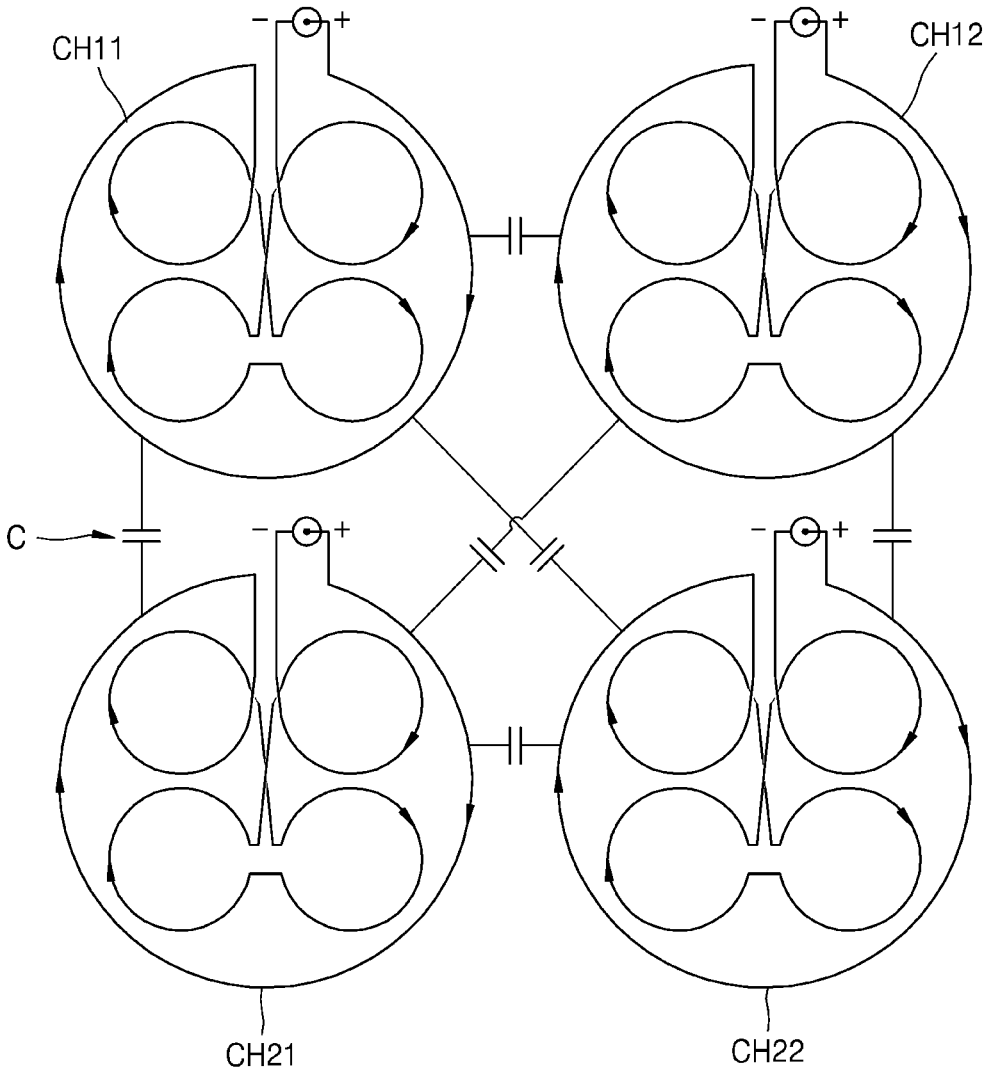[Fig 4]