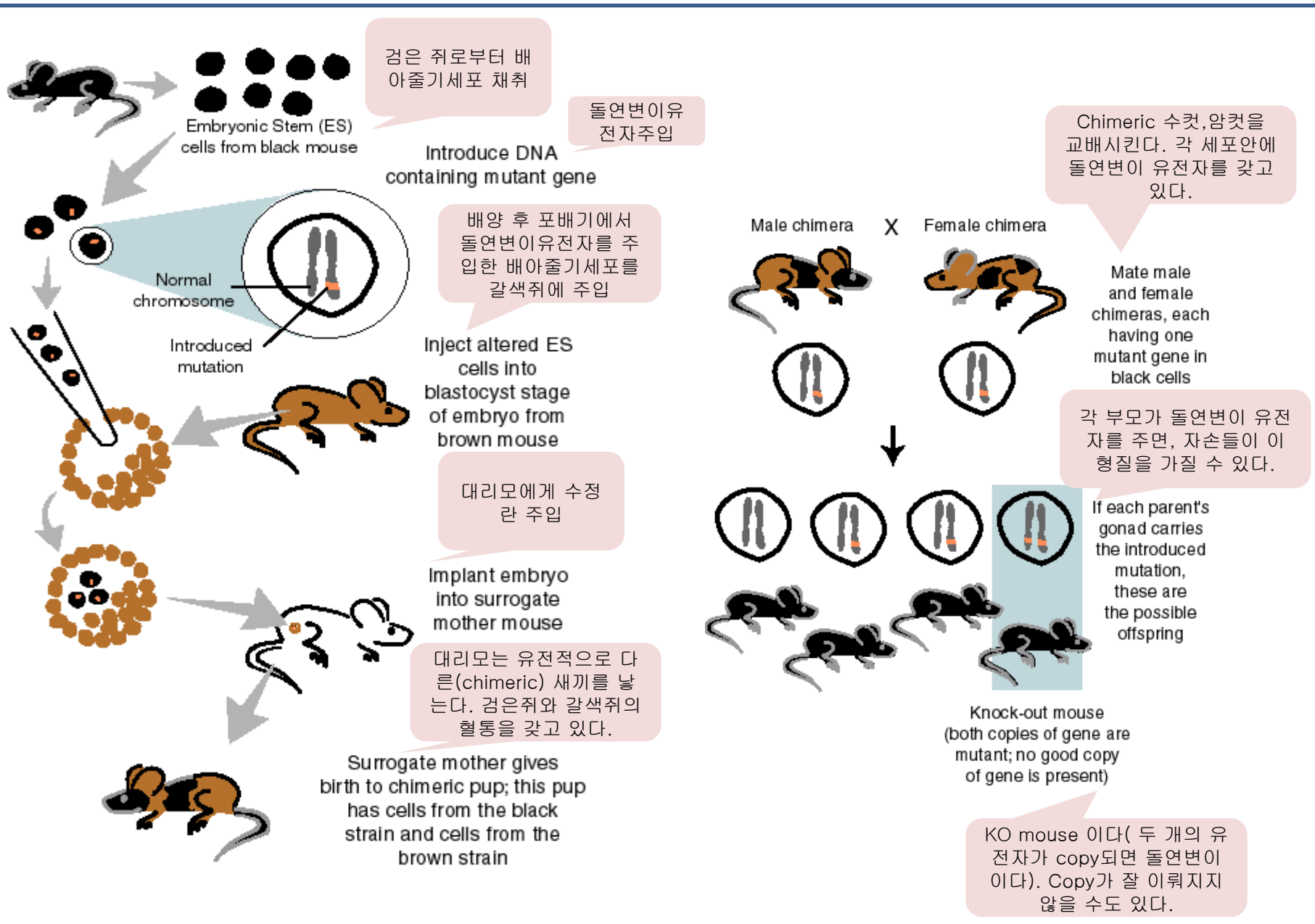Engineering Knock-Out Mouse