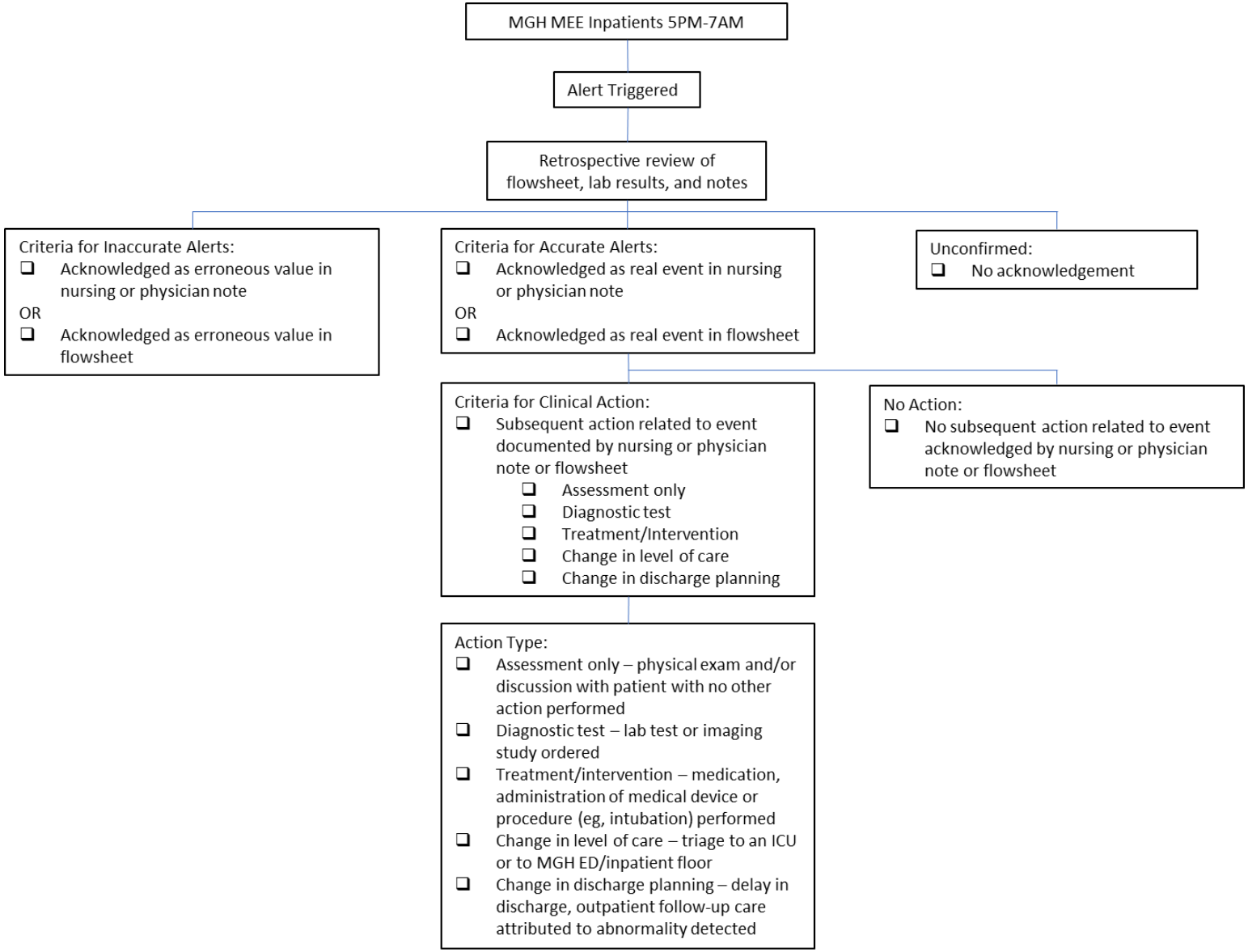Appendix Figure 1