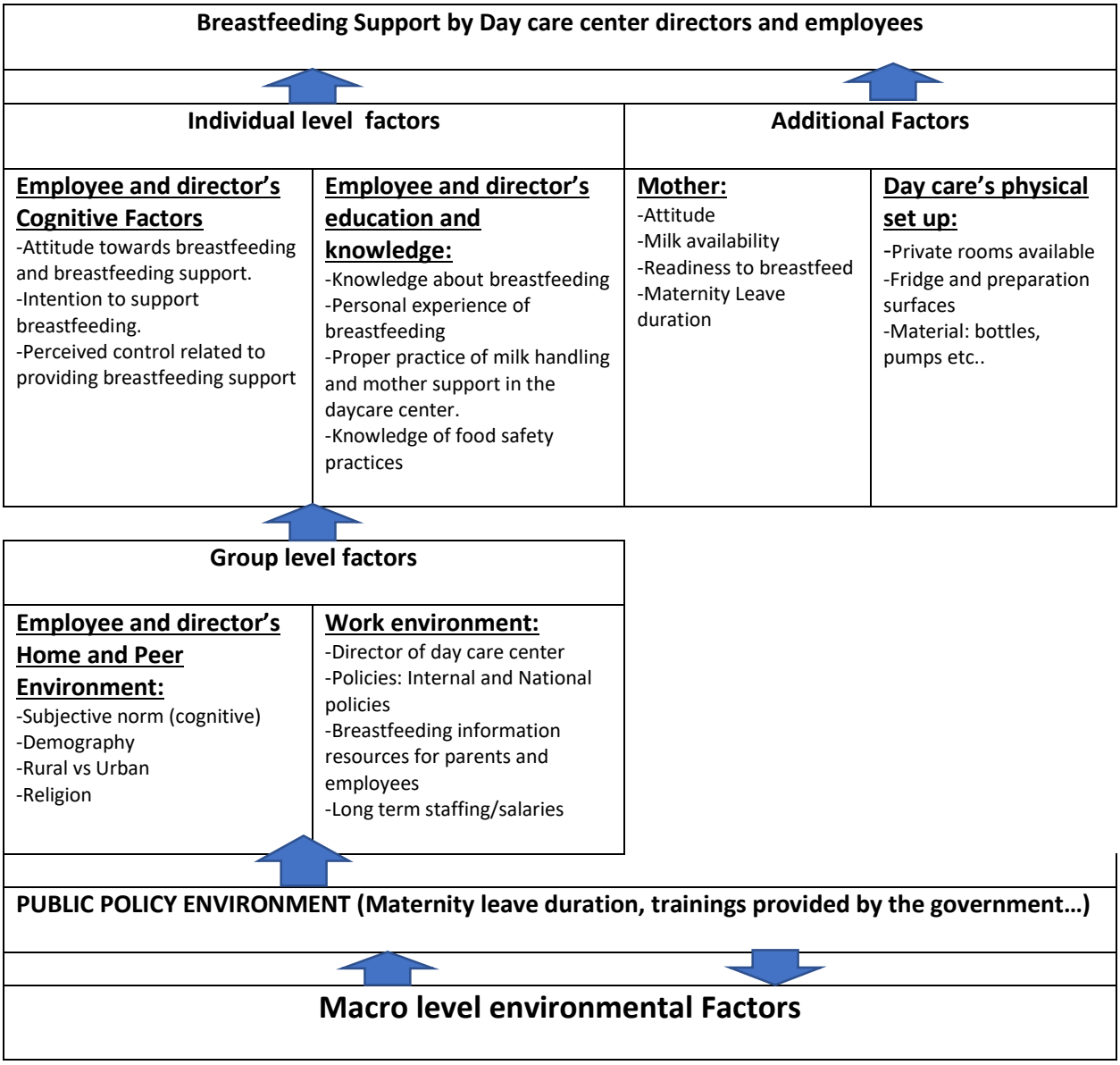Figure S1 Conceptual framework