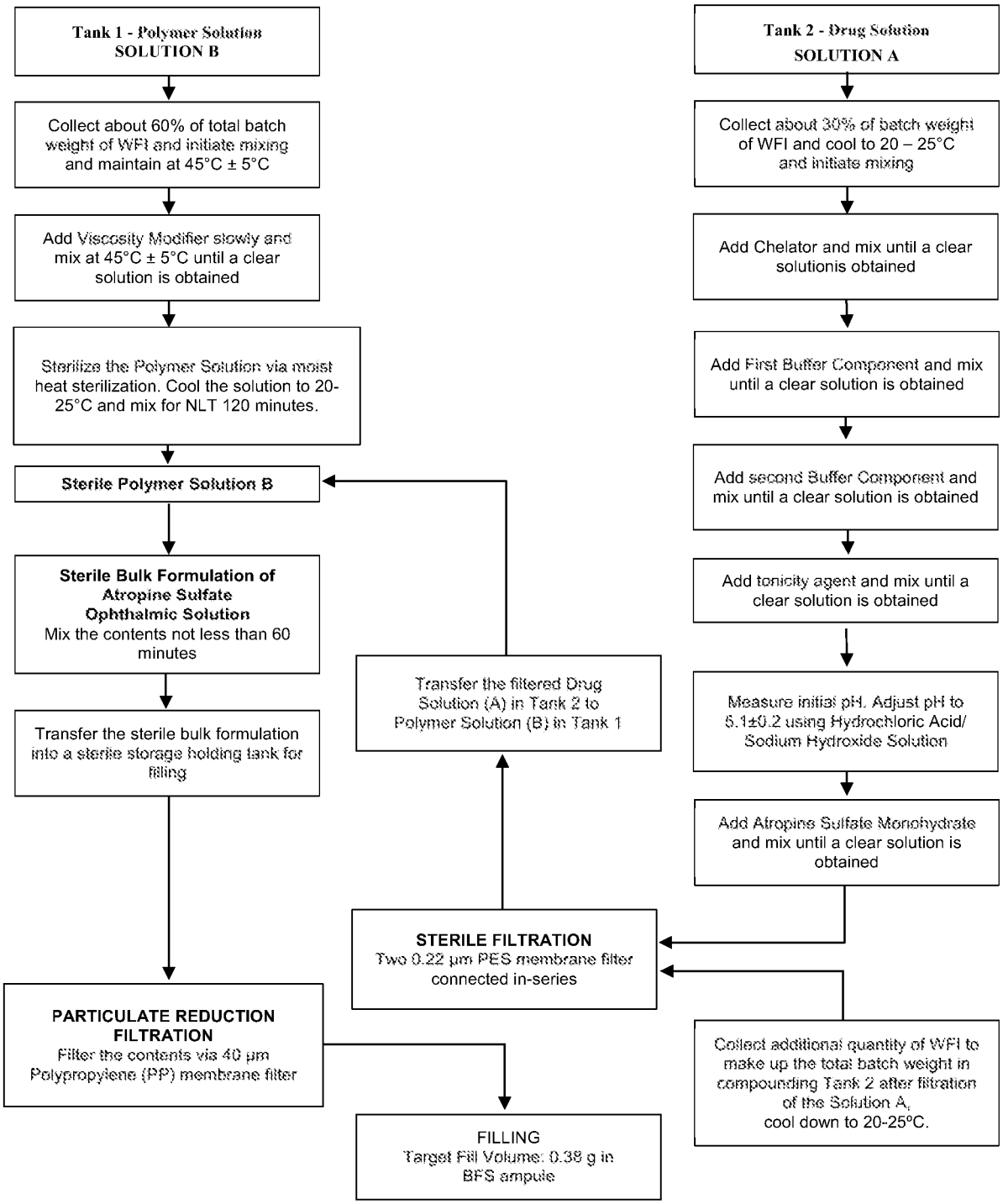

Tank 1 - Polymer Solution
SOLUTION B
Collect about 60% of total batch weight of WFI and initiate mixing and maintain at 45°C ± 5°C
Add Viscosity Modifier slowly and mix at 45°C ± 5°C until a clear solution is obtained
Sterilize the Polymer Solution via moist heat sterilization. Cool the solution to 20-25°C and mix for NLT 120 minutes.
Sterile Polymer Solution B
Sterile Bulk Formulation of Atropine Sulfate Ophthalmic Solution
Mix the contents not less than 60 minutes
Transfer the sterile bulk formulation into a sterile storage holding tank for filling
PARTICULATE REDUCTION FILTRATION
Filter the contents via 40 µm Polypropylene (PP) membrane filter
Tank 2 - Drug Solution
SOLUTION A
Collect about 30% of batch weight of WFI and cool to 20 – 25°C and initiate mixing
Add Chelator and mix until a clear solutionis obtained
Add First Buffer Component and mix until a clear solution is obtained
Add second Buffer Component and mix until a clear solution is obtained
Add tonicity agent and mix until a clear solution is obtained
Measure initial pH. Adjust pH to 5.1±0.2 using Hydrochloric Acid/ Sodium Hydroxide Solution
Add Atropine Sulfate Monohydrate and mix until a clear solution is obtained
Transfer the filtered Drug Solution (A) in Tank 2 to Polymer Solution (B) in Tank 1
STERILE FILTRATION
Two 0.22 µm PES membrane filter connected in-series
Collect additional quantity of WFI to make up the total batch weight in compounding Tank 2 after filtration of the Solution A, cool down to 20-25°C.
FILLING
Target Fill Volume: 0.38 g in BFS ampule

Figure 1