WORK PLAN AND METHODOLOGY

• Literature search
  - Literature search shall be carried out by following methodology
  - Journals of international repute Elsevier etc
  - STN, SciFinder search, Articles, Merck Index, Scopus, Text books etc
  - Web searches for the product and related information
  - Outsourced web article access through Scitex and Global document searches [Paid searches]

• Patent search & evaluation
  - Patent searches and evaluation shall be done by following methodology
  - USPTO, ESPACE NET web sites for US and EP patents
  - Paten lens for all other relevant patent searches
  - WIPO for PCT applications / Publication
  - Patent search & FTO document generation for strategy clearance

• Design of Formulation strategy
  - Design of strategy shall be primarily based on Patent searches
  - Shall be non infringing to all patents in the area
  - Strategy shall provide a pharmaceutically equivalent product to Aggrenox

• Procurement of actives, raw material, Packaging Material , RLD , Tools, equipments etc
  - Based on project needs all RM/ PM shall be procured from approved Quality sources
  - API of DMF quality shall be used for development , Packaging shall be novel & based on stability studies
  - RM shall be sourced from standard pharmacopoeial grade

• Preformulation studies
  - All studies with API & Innovator product
  - Complete Preformulation study as per ICH/ USFDA guideline wherever applicable
• **Development of analytical methods**
  - All methods shall be developed as per ICH guidelines
  - Methods of Assay, RS & Dissolution using HPLC methods
  - Development of Analytical methods for simultaneous release of acid from the formulation

• **Formulation development trials**
  - All development activities shall be carried out as per USFDA guidelines
  - Bench scale studies/ Lab scale studies shall be carried out to arrive upon a suitable composition & process
  - QBD/ DOE shall be used during development
  - ICH quality guidelines for stability studies
  - Development shall be done using GPCG 1.1 Wurster for Pelletization

• **Composition & Process optimisation**
  - This shall be done as per DOE & QBD USFDA recommendations
  - Process optimisation at GPCG 1.1 & Pam Glatt 125 equipments attached with bottom spray
  - Drug layering optimisation
  - Modified release Coating polymeric system optimisation

• **Dissolution method development & mapping**
  - All dissolution study shall be done using USP Type I & Type III methodologies
  - Comparison shall be done with Aggrenox® Reference product
  - In-Vitro mapping

• **Stability studies & Packaging evaluation**
  - Stability studies as per ICH recommendations
  - Packaging relative to Reference product

• **Scale up & Process evaluation**
  - Process optimisation shall be done on Pam Glatt 125 with Wurster attachment
  - Optimisation as per DOE if required
  - Effect of scale up on dissolution studies