
# Work Plan Schedule

| Task | Time | Goal | Outcome |
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| **C Thresholds** | | **In order to test the hypothesis that appropriate and scientifically justifiable thresholds exist, the Working Group must develop such thresholds. The development of appropriate thresholds will be the responsibility of the relevant sub-teams (Chemistry and Toxicology). Thresholds for Large Volume Parenterals, Small Volume Parenterals Prefilled Syringes and Ophthalmic dosage forms will be considered.** | |
| **C1** | **6/08** | The Working Group will agree on the outline of a process (or processes) designed to test the stated hypothesis by attempting to develop appropriate and scientifically justifiable thresholds related to leachables and extractables. | **Process Development** Outline of a process(es) designed to develop thresholds, and thereby test the hypothesis. |
| **C2** | **3Q08** | The WG will propose Threshold Models. Processes will proceed in parallel utilizing appropriate expertise from various Working Group members. | **Process Implementation** Safety and Analytical Threshold models will be proposed for each category. |
| **C2s1** | **3Q08** | The Toxicology sub-team will initiate development of appropriate SCT/QT for leachables, and provide guidance on safety considerations for extractables where appropriate. | **Strategies for Safety Concern Thresholds** Discuss best practice to asses thresholds, strategy forward etc (lack of available data) Gain an understanding for the application of thresholds for PODP. Explore a staged approach for leachable qualification |
| **C2s2a** | **3Q08** | The Chemistry sub-team will develop appropriate and scientifically justifiable analytical threshold models for extractables/leachables in the selected dosage forms. | **AET Models** Strategies developed for determining analytical threshold for extractables/leachables in the selected dosage forms. |

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| **C2s2b** | **4Q09** | The Toxicology sub-team will establish and implement the qualification/testing paradigm for leachables/extractables in PODP and propose the threshold values to be utilized.<br>. | **Development of SCT and QT**<br>A decision tree representing the qualification of leachables/extractables in PODPs will be proposed, thresholds for leachables/extractables in PODPs, including an example of a complete qualification for a representative PODP leachables profile. |
| **C3** | **1Q10** | The Working Group will thoroughly evaluate and conclude the results of the process implementation described under sub-tasks 2a and 2b and come to consensus as to the validity of the hypothesis based on the testing criteria previously stated. | **Harmonization and Consensus (Thresholds)**<br>A consensus within the WG of the thresholds and the successful completion of the mock qualification will be considered a successful test of the hypothesis. |
| **D<br>Best Practices** | | **Best demonstrated practices are the primary responsibility of the Chemistry sub-team. Actual testing and characterization of selected materials of common PODP container closure systems for each representative category will be evaluated with respect to extractables and subsequent correlation to theoretical leachables. In order to test the hypothesis that best demonstrated practices for the characterization and analytical evaluation exist, the Working Group will establish such practices.** | |
| **D1** | **3Q08** | The Chemistry Team will develop the Test Protocol, defining the objectives of the study, the materials to be characterized, the contact media to be used, the contact conditions and the test methods. | **Test Protocol Development**<br>Implementable Test Protocol(s) shall be developed for each category of the selected dosage forms and approved by all Working Group members. |
| **D2** | **4Q08** | The Chemistry sub-team will develop the Test Plan, which details the mechanics and timing for the implementation the Test Protocol(s). Activities associated are as follows:<br>Identification and recruitment of test material and test system suppliers.<br>Identification and recruitment of auxiliary team members.<br>Identification of Study Leaders<br>Delineation of study responsibilities (Deliverables and Schedule).<br>Development and approval of the Test Plan and Schedule<br>Procurement of the Test Materials.<br>Generation of the Test System(s).<br>Distribution of the Test Materials and Test System(s) to the auxiliary team members. | **Test Plan Development**<br>The Test Plan will be approved by the Chemistry sub-team and all auxiliary team members. Auxiliary team members will be provided with appropriate test materials and systems. At this point, the Chemistry sub-team will be ready to implement the Test Plan. |

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| **D3** | **3Q09** | The auxiliary team members shall perform the required testing and provide results to the Study Leader. | **Test Plan Implementation**<br>Testing will be completed and Test Results communicated to the Study Leaders. The results for each category will be summarized and reported as available to PQRI. |
| **D4** | **2Q10** | The Study Leaders shall review and collate the test results from the study participants and draft a comprehensive Study Report will be produced, which will include results from all categories, evaluated and demonstrated practices.<br>Members of the Chemistry sub-team will review the draft Study Report and provide appropriate feedback.<br>The Chair of the Chemistry sub-team shall collate feedback, amend report, and provide the Working Group and PQRI with a Final draft of the Study Report. | **Identification of Best Demonstrated Practices**<br>Final Draft of Study Report for Working Group and PQRI review. |
| **D5** | **2Q10** | The Working Group will thoroughly evaluate the results of the process implementation described under Tasks D3 and D4 and come to consensus as to the validity of the hypothesis based on the testing criteria previously stated. | **Harmonization and Consensus (Best Practices)**<br>The Working Group as a whole will critically evaluate the outcomes of Tasks D3 and D4 and create a comprehensive thresholds and best practice report for review within the PQRI process that will include all proposed outcomes as well as clearly stated recommendations for the technical community, including points for the Agency (FDA) to consider. |
| **A consensus within the Working Group on thresholds/best demonstrated practices and the completion of Tasks C&D will be considered a successful test of the hypothesis. The expected deliverable date is estimated 2Q10** | | | |
| **Other** | | Other outcomes may include publications and presentations at appropriate scientific meetings and forums. . | These additional outcomes will be discussed and agreed to at the appropriate time in the overall PQRI process |