The mission of the CDISC Integrated Safety Data Pilot is to demonstrate that an integrated data submission (including analysis plan and key analyses) created using the CDISC Standards is technically feasible and will meet the needs and expectations of scientific reviewers in conducting an integrated patient and population review of data from multiple studies and compounds.
CDISC Pilot ISD Goals

- Assess the applicability of the CDISC Standard to integrate data
- Identify any issues/questions to be addressed by the CDISC Standards Teams
- Validate the components of the CDISC standard can effectively be used together
CDISC Pilot ISD Goals

- Use the CDISC models to convert legacy safety data into an integrated database
- Use the CDISC models to support integrated standard safety analysis and reporting as described by the current FDA Guidances
- Evaluate the most current CDISC models including SDTM, ADaM, ODM/Define.xml, and Trial Design
- Support the critical path initiatives around standard data, integrated databases, standard data collection, and studies of special populations
Overview of Pilot Data

- 3 compounds
  - Multiple Clinical trials
  - Multiple pharmacological classes

- Trial Designs
  - Pharmacokinetics trials
  - Randomized double-blind trials

- Artificial test data composed of deidentified and randomly modified data; will be provided as SDTM
  - “Practice” Data
Overview of Pilot Data

- Documentation provided:
  - One page summary of each study including study design schedule
  - Need to determine how to generate annotated CRFs with the SDTM fields

- Initial domains included: DM, AE, VS, EX, CM, DS, SC, PC, LB, MH
Decisions (preliminary)

- Pilot will use the most current models (draft or final)
- Submission will not be a traditional ISS but a safety profile
- Derived data in SDTM will be limited
- ADAM will be generated both for each study and the integration
- SDTM will only be generated for each study

Of course all decisions are subject to change
Analysis/Design Update

- Three compounds
  - Compound A - 4 studies
  - Compound B - 2 studies
  - Compound C - 2 studies
Goal is to deliver a profile and not a safety assessment

Three levels of design and analysis
- Primary-Individual study level with option to use design model to generate population profile
- Secondary-Compound level where only populations that had exposure to a specific compound will be included and integrated
- Tertiary-Disease level where population profiles from all compounds are integrated
Analysis/Design Update

- **General Plan**
  - Adverse events are both clinical observations and abnormal laboratory values
  - Clinical observations will be flagged as serious and graded when feasible
  - Laboratory observations will be flagged as abnormal and graded when feasible
  - Concomitant medications will not be included in analysis
  - Medical history for events occurring prior to study enrollment will not be included in analysis
Analysis/Design Update

- Output will be composite frequency listing of adverse events and abnormal laboratory observations
- Listings will be overall and by serious and grade if available
- Additional analyses will be by exposure level for each compound, age (<2 years, 3-5 years, 6-10 years and 11+), and gender
Data/Programming Update

- Summary of domain review
- Update on metadata collection
- Overview of programming process
- Questions and next steps
Summary of domain review

- SAS provided raw datasets
- Domains include DM, SC, MH, CM, EX, LB, VS, AE, DS and PC
- Documentation includes Study_Overview and Domain Summary
- Team members were assigned to domains to compared the Raw vs SDTM IG 3.1.1
- Team members were assigned to compare SDTM IG 3.1.1 vs 3.1.2 (draft)
Domain review - assignments

- DM, SC
- AE
- VS
- EX
- DS
- LB
- PC
- MH, CM – under review in inclusion for analysis
Domain review - assignments

- Controlled terminology review – compare raw vs published –
- Pilot domain review – IG 3.1.1 vs IG 3.1.2 –
Submission deliverables

- For each study:
  - SDTM datasets, define.xml
  - ADaM datasets, define.xml
  - Study report with tables, listings and graphs

- For Integrated database (by compound and overall for all compounds):
  - ADaM datasets, define.xml
  - Integrated report with tables, listings and graphs
Next Steps

- SAPs completed
- Programming underway
- Continue the project work on Programming and Package
- Team Leaders meet weekly project map updated
- Multiple customers
- Projected delivery 4th quarter 2008