Medical **D**ictionary for **R**egulatory **A**ctivities (MedDRA) Update

International Conference on Harmonisation of Technical Requirements for Registration of Pharmaceuticals for Human Use

Topics

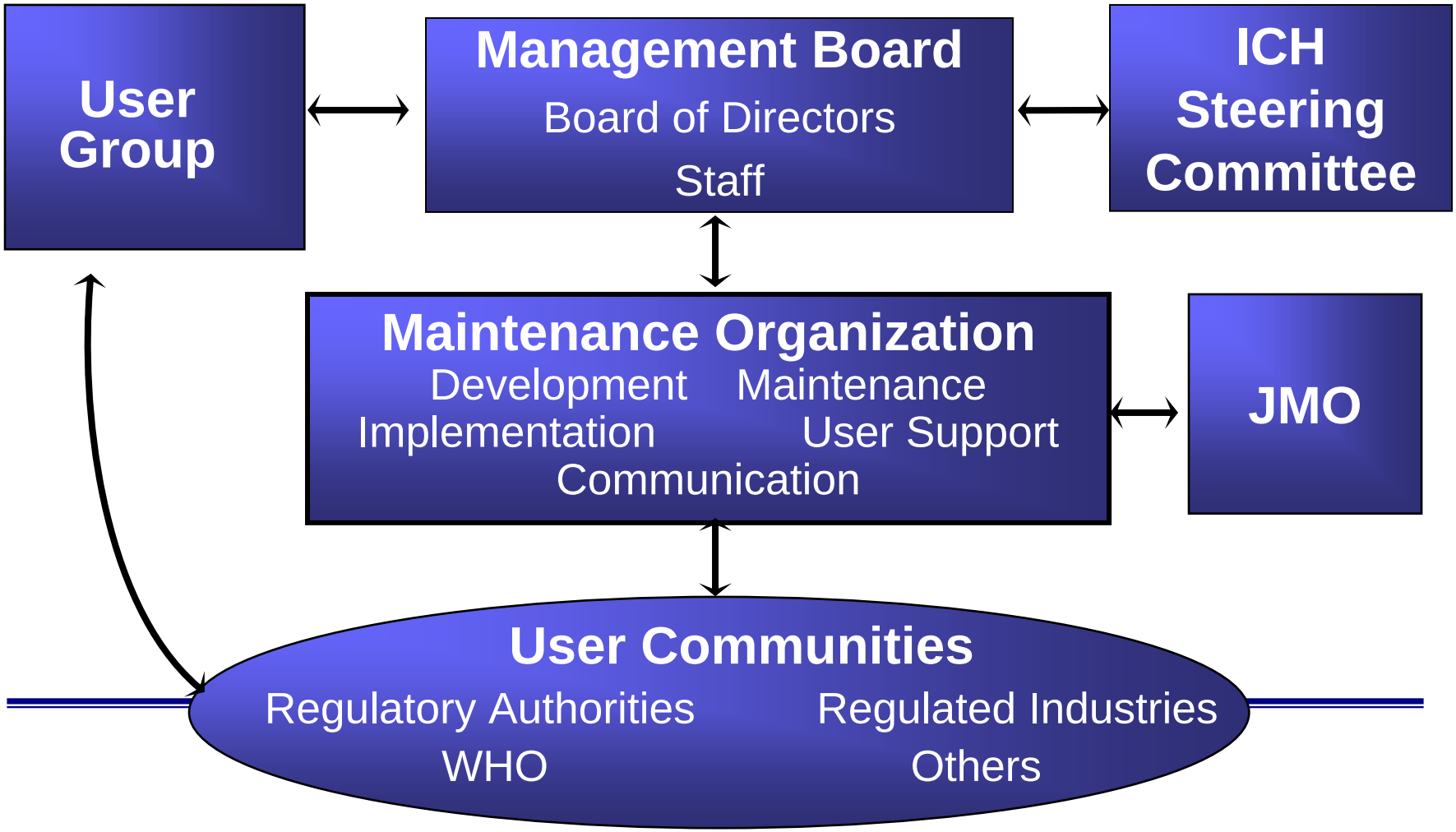
- MedDRA as an ICH initiative
- How MedDRA is used
- Standardised MedDRA Queries (SMQs)
- Recent developments
  - MedDRA and Vigibase
  - Chinese MedDRA Translation
  - CTCAE Developments
  - Medical Device Terms Developments
  - MedDRA Free Training

Objectives for MedDRA Development

Result of an ICH initiative (M1)

- An international multi-lingual terminology
- Standardized communication between industry and regulators
- Support of electronic submissions
- Application through all phases of the development cycle
- Classification a wide range of clinical information for multiple medical product areas
- Maintained by MSSO

Maintenance Organization Structure



How is MedDRA Used?

- To code (classify) ADR/AEs, history, indication, investigations, procedures, etc.
- Verbatim/reported information is linked to a MedDRA Lowest Level Term (LLT)
- LLTs have a parent concept Preferred Term (PT)
- Grouping terms and System Organ Classes (SOCs) logically group concepts for later retrieval and analysis
- Used for coding, assessment, and reporting of safety data for both marketed products and clinical studies

MedDRA Hierarchy

- SOC Hepatobiliary disorders
- SOC Immune system disorders
  - HLGT Allergic conditions
    - HLT Allergic conditions NEC
    - HLT Allergies to foods, food additives, drugs and other chemicals
    - HLT Anaphylactic responses
      - PT Anaphylactic reaction
      - PT Anaphylactic shock
      - PT Anaphylactic transfusion reaction
      - PT Anaphylactoid reaction
        - LLT Anaphylactic type reaction
        - LLT Anaphylactoid reaction
        - LLT Delayed anaphylactoid reaction
      - PT Anaphylactoid shock
      - PT Anaphylactoid syndrome of pregnancy
      - PT First use syndrome
    - HLT Angioedemas
    - HLT Atopic disorders
    - HLT Urticarias
  - HLGT Autoimmune disorders
  - HLGT Immune disorders NEC
  - HLGT Immunodeficiency syndromes
- SOC Infections and infestations

Definition of SMQ

- Result of cooperative effort between CIOMS and ICH (MSSO)
- Groupings of terms from one or more MedDRA System Organ Classes (SOCs) related to defined medical condition or area of interest
- Included terms may relate to signs, symptoms, diagnoses, syndromes, physical findings, laboratory and other physiologic test data, etc., related to medical condition or area of interest
- Intended to aid in case identification

SMQs in Production - Examples

- As of Version 11.1, a total of 67 in production (Other SMQs in development)

- Adverse pregnancy outcome/reproductive toxicity (incl neonatal disorders)
- Agranulocytosis
- Anaphylactic reaction
- Cerebrovascular disorders
- Convulsions
- Depression and suicide/self-injury
- Hepatic disorders
- Ischaemic heart disease
- Lack of efficacy/effect
- Peripheral neuropathy
- Pseudomembranous colitis
- Rhabdomyolysis/myopathy
- Severe cutaneous adverse reactions
- Systemic lupus erythematosus

MedDRA in Vigibase

- MSSO has worked with WHO at the Uppsala Monitoring Center to include MedDRA in Vigibase
- Vigibase includes several million AE cases collected around the world
- All Vigibase events now have a MedDRA term assigned
- Vigibase provides searches and reports that support MedDRA
- Became available in April 2008

Chinese MedDRA – Current Status

- Currently conducting the translation
  - Will be maintained like all other translations
- MSSO will develop a webpage on MSSO website in Chinese to provide basic facts
- Anticipate initial release in late 2009
- Supported by updated MedDRA Desktop Browser
- Fee to be on a sliding scale

CTCAE and MedDRA

- Common Terminology Criteria for Adverse Events v3.0,required by NCI to code trials
- Sponsors often code in both CTCAE and MedDRA
- Current CTCAE/MedDRA mapping insufficient
- NCI to include MedDRA LLTs directly into CTCAE
  - New version (4.0) will be available in April 2009
  - Removes need for double coding or map to MedDRA
- MSSO working closely with NCI and cancer community to develop revision

Medical Device Terms

- CDRH
  - 60 terms already added to MedDRA
- ISO (TC 210 WG)
  - “Coding structure for adverse event type and cause”
  - Very general terms
- MSSO Device Working Group tasks
  - Industry, FDA, JMO
  - Review CDRH and ISO terms
  - Develop recommendations on criteria for inclusion
  - Develop recommendations on the future device hierarchical structure
  - Expected to take 6 months

Free MedDRA Training (cont)

- Training came as a result of concerns from regulatory authorities
  - Poor coding quality
  - Turnover in industry
  - Need to rely on electronic data exchange
- MSSO is conducting free MedDRA training classes
  - Coding with MedDRA
  - Introduction to MedDRA Data Analysis and SMQs for Physicians
- Conducted in the US and Europe
  - 35 classes scheduled, 22 completed
- Free webinars being scheduled

Questions?