How to handle study drug (investigational drug)

Investigational Product

- A pharmaceutical form of an active ingredient or placebo being tested or used as a reference in a clinical trial, including
  1. An investigational new drug
  2. A marketing approved drug when
     - has a new dosage form
     - use for an unapproved indication
     - need to gain additional information

ICH GCP 1.33

Labeling

- Label should include a statement that the product is limited to investigational use
- No false or misleading statement
- Understandable to subjects
- Avoid misuse or medication error

ยาใช้เพื่อการศึกษาวิจัยเท่านั้น

RESPONSIBILITIES

Investigator's Responsibilities

1. Has duty for the study drug accountability
   - May assign the duty to an appropriately qualified site staff
2. Maintain records of and reconcile the study drug that
   - delivered to and inventoried at the site
   - used by each subject
   - returned to the sponsor or other alternative disposition

(ICH E6, 4.6.1 – 4.6.6)

Investigator's Responsibilities

4. Store the study drug as specified by the sponsor
5. Use the study drug only in accordance with the approved protocol
6. Explain the correct use of the study drug to each subject
7. Check that each subject is following the instructions properly, at interval appropriate to the study

(ICH E6, 4.6.1 – 4.6.6)

Sponsor's Responsibilities

1. Study drug or placebo is manufactured in accordance with any applicable GMP
2. Labeling comply with applicable regulatory requirement
3. Determine storage temperatures, storage conditions (e.g. protection from light), storage time of the study drug
4. Determine administration procedures
5. Should not supply an investigator with the study drug until obtaining approval from IRB/IEC and regulatory authority (if applicable)

Sponsor's Responsibilities

6. Ensure timely delivery of study drug to the investigators
7. Maintain records that document shipment, receipt, disposition, return, and destruction of the study drug
8. Maintain a system for retrieving study drugs and documenting this retrieval)
9. Maintain a system for the disposition of unused study drug and for the documentation of this disposition

Sponsor's Responsibilities

10. Ensure that the study drug is stable over the period of use.
11. Maintain sufficient quantities of the study drug
12. Maintain records of batch sample analyses and characteristics
13. Retain samples until the study data analysis is complete or as required by applicable regulatory requirement
14. Instructions for the handling and storage of study drug

HANDLING OF INVESTIGATIONAL DRUG

Receiving Investigational Drug Shipments

- Once receive the investigation drug:
  1. Check the shipment and confirm that the drug content and label are intact
  2. No excursion of temperature record during shipment (if applicable)
  3. Compare the invoice to the batch/lot no., expiry date, quantity and dosage, code no.
  4. Report the sponsor immediately for any discrepancy of record, drug/label damage or temperature excursion

Storing Investigational Drug

- Investigational Drug Storage Procedures:
  1. Separate from the other medicines
  2. Temperature control
     - Thermometer validation
     - Temperature record (temperature log)
     - Backup system
     - Handling of temperature excursion
  3. Secure storage/restricting accessibility
     - Access control (securely lock)
     - Access record

Dispensing Investigational Drug

1. Assign appropriately qualified person and document the assignment
2. Adhere to randomization process
3. Dispense the study drug only to the assigned subject with explanation of use
4. Prepare, administer, dispense the study drug in accordance with the approved protocol including dosing, dose modification and discontinuation
5. Adhere to blinding procedure (especially when unblinded pharmacist required)

Dispensing Investigational Drug

6. Do not change study drug from one subject to another
7. Check inventory of study drug and monitor expiry date
8. Inform the sponsor once the study drug is near expiration or short of
9. Do not substitute study drug from hospital stock

Returning Investigational Drug

1. Collect containers and unused study drug
2. Check that each subject is following the instructions properly
3. Reconcile drug accountability record and return all unused and unopened drug to sponsor (including empty containers)

How to Record Study Drug Received, Dispensed, Returned

Recording Drug Accountability

1. Drug receipt: from the sponsor to site
2. Subject dispensing from site to each subject at each study visit
3. Subject return: from each subject to site at each study visit to sponsor or destruction
4. Drug return: from site to the sponsor for used (container), unused, unopened (not dispended) drug
5. Drug Destruction

Drug Accountability Record

- Treatment assignment and dose adjustment (in source document) as specified in the protocol
- Missing/lost or damage study drug
- Reconcile all study drugs

ยารับมา - ยาผู้ป่วยใช้ - (ยาที่ทำหายไป) = ยาที่คืน

Record of Drug Receipt and Return

1. Date received
2. Drug name
   - Dosage form (tablet, capsule etc.) and strength; package (bottle, blister pack, vial etc.)
   - Batch no. or manufacturing lot no.
   - Expiry date
   - Code no. (randomization code, subject code); dispensing visit; quantity received
3. Date returned
4. Quantity returned
   - Balance of quantity dispensed and return (used, unused, lost/damage) and unopened (not dispensed)
5. Signature of assigned person

Record of Subject dispensing and Return

1. Subject ID code
2. Study visit/date dispensed
3. Drug name
   - Dosage form and strength
   - Batch no. or manufacturing lot no.
   - Expiry date
   - Code no. (randomization code, subject code); dispensing visit; quantity dispensed
4. Date returned
5. Quantity returned (used, unused, lost/damage)
   - Balance of dispensed drug and returned drug
6. Signature of assigned person

Key things to remember

- Complete drug accountability records in a timely manner
- Periodically verify and reconcile drug accountability records
- Keep drug in appropriate temperature and secure place
- Observe drug expiry date and maintain adequate supply
- Document drug destruction

QUESTION & ANSWER