1st Thailand GRM Conference, Bangkok, Thailand, June 26-28, 2018

Basic Concept of Good Registration Management

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http://www.fda.gov.tw/

Outline

- Concept of Good Registration Management (GRM)
- APEC GRM Roadmap with its key outcomes
- Training programs for APEC
- Performance indicators
- Summary

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- **Concept of Good Registration Management (GRM)**
- APEC GRM Roadmap with its key outcomes
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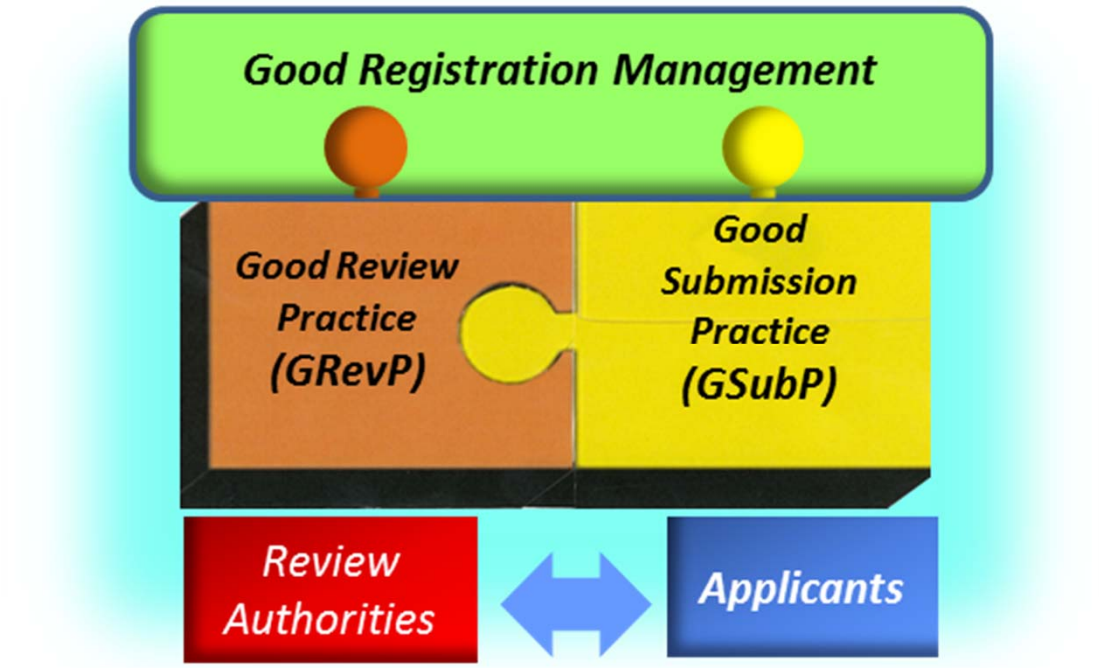
Concept of Good Registration Management



GRM: *A concept to promote efficient registration process for medical products by promoting Good Review Practice (GRevP) and Good Submission Practice (GSubP) cooperatively.*

Goal of GRM: *To benefit the patients with timely access of medical products of safe, efficacious and good quality*

Definitions and Objectives (1)

Good Review Practice

Definition

GRevPs are documented best practices for any aspect related to the process, format, content and management of a medical product review.

Objective

The objective is to help achieve timeliness, predictability, consistency, transparency, clarity, efficiency and high quality in both the content and management of reviews.

Definitions and Objectives (2)

Good Submission Practice

Definition

An industry practice for any aspect related to the process, format, contents and management of submission for registration of medical products by applicants.

It is the practice to enhance the quality and efficiency of the product registration process by improving the quality of submission as well as its management.

Objective

The objective is to help applicants prepare good quality submission leading to successful registration

Outline

- Concept of GRM and its objective and goal
- **APEC GRM Roadmap with its key outcomes**
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APEC Roadmap to Promote GRM (2011-2020)

Co-champions: Chinese Taipei and Japan

Goals of the GRM Roadmap

To **promote the concept of GRM** and thereby enhance mutual trust for regulatory convergence among the APEC member economies by 2020

Goals of the key elements of GRM

Good Review Practice	Good Submission Practice
To **strengthen the performance, predictability, and transparency of regulatory agencies** through the implementation or enhancement of GRevP and quality measures stepwise in each interested APEC economy	To **enhance the quality and efficiency of the medical product registration process** by improving the quality of submission as well as its management

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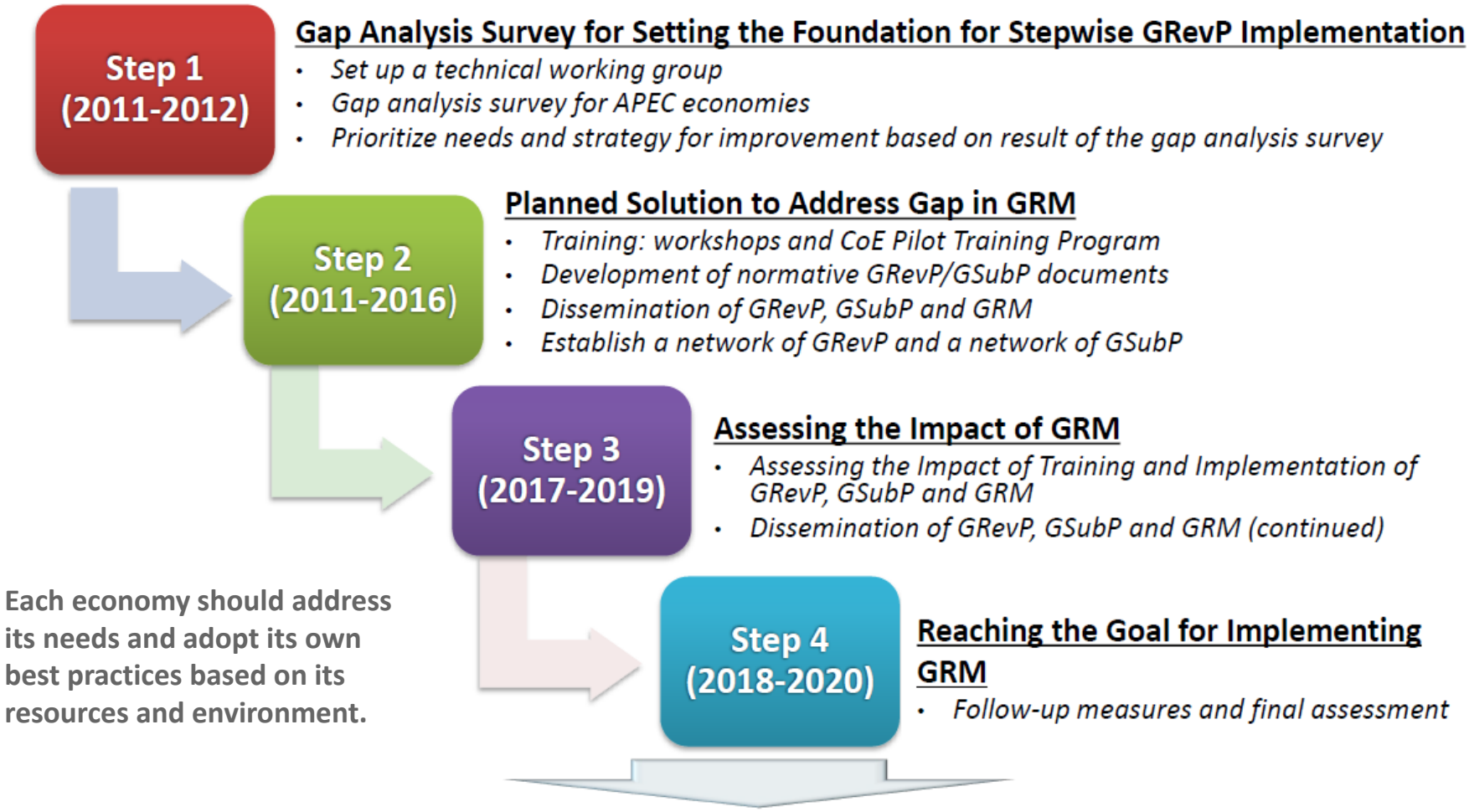
Regulatory Convergence

RHSC definition

- **Regulatory convergence** represents **a voluntary process** whereby the regulatory requirements across economies **become more similar or "aligned" over time** as a result of the gradual adoption of **internationally recognized** technical guidance documents, standards and scientific principles (harmonization) and **common or similar practices and procedures**.
- It does not represent the harmonization of laws and regulations, which is not necessary to allow for the alignment of technical requirements and for greater regulatory cooperation.

Specific Activities and Timeframe of the GRM Roadmap



Key Outcomes from Step 1

Step 1	Step 2	Step 3	Step 4
2011-2012	**2011-2016**	**2017-2019**	**2018-2020**
Gap Analysis Survey for setting the foundation for Stepwise GRevP Implementation	Planned solutions to address gaps GRM CoE Pilot	Assessing impact of GRM using Performance indicators	Reaching the Goal for Implementing GRM

Characterizing Good Review Practices: A Survey Report Among Agencies of APEC Member Economies

Gap analysis published (2013)

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Gap analysis survey for good review practices

Gap Analysis for GRevP in APEC	• **CIRS** conducted a gap analysis survey among regulatory agencies of **14 APEC member economies** in **2011-2012**. • **Questions:** agency characterization, good review practices, timeliness, transparency, quality, predictability
Status of GRevP implementation in 2012	• The majority established GRevP. Most practice are evolving and are applied on an **informal basis**. • Most agencies have developed **SOPs and guidelines** and **use a variety of training methods**.
Identified training needs	• Use of **assessment templates or frameworks** • **Implementation of GRevP** • How to **develop and improve SOP**
Advantages of a common review process	• Build **trust and confidence** in agency's processes • Setting the stage for the possibility of **work sharing** • Bringing **consistency** and **transparency** to review process

Liu L-L et al. Therapeutic Innovation & Regulatory Science, November 2013; vol. 47, 6: pp. 678-683.

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Key Outcomes from Step 2

Step 1 | **Step 2** | **Step 3** | **Step 4**

2011-2012

Gap Analysis Survey for setting the foundation for Stepwise GRevP Implementation

2011 APEC GRevP Workshop

2012 APEC GRevP Workshop

2011-2016

Planned solutions to address gaps

GRM CoE Pilot

GSubP PWA endorsed (2014)

GRevP Guidelines (2015)

GSubP Guidelines (2016)

GRM Roadmap endorsed (2016)

2017-2019

Assessing impact of GRM using Performance indicators

2018-2020

Reaching the Goal for Implementing GRM

APEC GRM Training Activities

1. TFDA/RAPS (CoE)
- 2016 pilot (Nov 2016, Taipei)
- 2017 workshop (Oct 2017, Taipei)
- 2018 workshop (Sep 2018, Taipei)

2. COFEPPRIS (pilot CoE)
- 2017 pilot (Jun 2017, Mexico City)

3. Local Training
- 2017: Singapore
- 2018: Taiwan, Thailand, Malaysia

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Topics Discussed in the 2011 and 2012 GRevP Workshops

2011 Basic Workshop	2012 Advanced Workshop
A. The Basics 1. Common Understanding of GRevPs 2. Industry's role, responsibilities B. The Details 1. Reviewer Competencies 2. Reviewer Training 3. Templates and Procedures C. Metrics D. Information Resources 1. Use of Peer Review 2. Use of External Experts 3. Transparency & Information Sharing E. Recommendations and Final Remarks F. Good Review Practices on Medical Products G. Industry Responses H. Panel Discussion	A. Review of 2011 APEC Basic GRevP Workshop B. Quality System for Reviewers C. Key Elements & Strategies of a Good Review D. Critical Thinking and Decision Making: Drugs and Devices E. Transparency and Interactions: With the Public, Industry/Other Stakeholders and Regulatory Authorities F. Conclusion

- Lin H-Y et al. Therapeutic Innovation & Regulatory Science, November 2015; vol. 49, 4: pp. 483-492.
- https://www.fda.gov.tw/EN/site.aspx?sid=3309

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Literature survey for good submission practices

- **GSubP:** Several articles addressed the issue of quality of application submissions
  - Indicate necessity of promotion of **GSubP by applicants** & **GRevP by regulatory authorities**

- Independent Evaluation of FDA's First Cycle Review Performance – Retrospective Analysis Final Report. January 2006 (by Booz Allen Hamilton Inc.)
- Characterizing Good Review Practices: A Survey Report Among Agencies of APEC Member Economies. TIRS 47(6) 678-683, 2013
- Building Quality into Regulatory Activities: What does it mean? June 2006 (by CMR International

Endorsement of GRevP and GSubP Guidelines

GRevP Guidelines (WHO, 2015)

Table of Contents

1. Introduction
2. Glossary
3. **Principles of a good review**
4. **Managing the review**
5. **Communications**
6. **Review personnel**
7. **Conducting the review**

Bibliography

http://apps.who.int/medicinedocs/en/d/Js21902en/

GSubP Guidelines (APEC RHSC, 2016)

Table of Contents

1. INTRODUCTION
2. **PRINCIPLES OF A GOOD SUBMISSION**
3. **MANAGEMENT OF SUBMISSION**
4. **COMMUNICATIONS**
5. **COMPETENCY AND TRAINING**
6. GLOSSARY
7. REFERENCE

http://apac-asia.com/images/achievements/pdf/5th/2_APEC_RHSC%20Endorsed%20GSubP%20Guideline.pdf

These 2 documents serve as:

- Guidelines for regulatory authorities conducting application review and industries conducting application submission
- Basis for the APEC GRM Roadmap champions to develop training objectives and core curriculum

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Training Objectives

Principles

- The principles of Good Review Practices (GRevP) and Good Submission Practices (GSubP)

Good Review

- What is needed for regulators to accomplish good review
  - Conducting and managing the review
  - Good communication with applicants
  - Competency for regulators

Good Submission

- What is needed for applicants to accomplish good application
  - Planning and preparation of application dossier
  - Good communication with regulators
  - Competency for applicants

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Core Curriculum

GRM
Good Registration Management

Common Sessions

- Basic concept of GRM
- An Overview of Good Review
- An Overview of Good Submission
- Effective Communication for GRM
- Competency & training
- Rolling out the GRM training program in each economy

GRevP
Good Review Practices

Reviewers-Specific Sessions

- Managing the review - an Overview
- Communication : Fundamentals and Case Studies
- Review personnel - Critical thinking
- Conducting the review

GSubP
Good Submission Practices

Applicants-Specific Sessions

- Planning of Application
- Preparation of application dossier / Practice : How to prepare application dossier
- Effective communications Focusing follow-up actions during review period

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Hosting 2016 pilot workshop and endorsement of full CoE

Date: November 15-17, 2016

Venue: Chang Yung-Fa Foundation, Taipei

Co-organizers:

Participating economies (17): Chile, China, Hong Kong China, Indonesia, Japan, Korea, Malaysia, Mexico, Papua New Guinea, Peru, Philippines, Singapore, Chinese Taipei, Thailand, Viet Nam, USA, and UK

Endorsement of full CoE: In February 2017, TFDA/RAPS were endorsed as a formal CoE by APEC LSIF-RHSC.

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Program of 2016 GRM CoE Pilot Workshop

<table>
<tr><th>Day 1</th><th colspan="2">Day 2</th><th colspan="2">Day 3</th></tr>
<tr>
<td rowspan="2">Common Sessions

Basic concept of GRM

An Overview of Good Review

An Overview of Good Submission

Case study: Effective Communication for GRM</td>
<td rowspan="2">Reviewer Sessions

Managing the review – an Overview

Communication: Fundamentals and Case studies</td>
<td rowspan="2">Applicant Sessions

Planning of Application

Preparation of applicant dossier/ Practice: How to prepare application dossier</td>
<td>Reviewer Sessions

Review personnel – Critical thinking

Conducting the review

Rolling out the GRM training program in each economy</td>
<td>Applicant Sessions

Effective communications - Focusing follow-up actions during review period

Rolling out the GRM training program in each economy</td>
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<tr>
<td colspan="2">Common Session

Panel discussion: Competency and Training</td>
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http://www.raps-in-taiwan.org.tw/apec/index.html

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Hosting 2017 APEC GRM CoE Workshop

Date: October 30 – November 2, 2017

Venue: NTU Hospital International Convention Center, Taipei

Co-organizers:

Participating economies (14): Canada, Hong Kong China, Indonesia, Japan, Korea, Malaysia, Papua New Guinea, Philippines, Singapore, Chinese Taipei, Thailand, Viet Nam, USA, and UK

2017 APEC GRM CoE Workshop Photos

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Program of 2017 GRM CoE Workshop

<table>
<tr><th>Day 1</th><th colspan="2">Day 2</th><th>Day 3</th></tr>
<tr><td colspan="2"><u>Common Sessions</u>

Keynote speech: Basic Concept of GRM

Overview of Good Review/ Submission

Experience sharing from different APEC member economies</td><td rowspan="2"><u>Reviewer Sessions</u>

Review personnel – Critical thinking (new drug vs. generic drug applications)

Communication: Fundamentals & Case studies</td><td rowspan="2"><u>Applicant Sessions</u>

Preparation of application dossiers

Communication during review period</td><td rowspan="2"><u>Common Sessions</u>

Communication
-Practices and interactive discussions between reviewers and applicants

Competencies and training for reviewers and applicants

Rolling out the GRM in each economy</td></tr>
<tr><td><u>Reviewer Sessions</u>

Managing & Conducting the review (new drug vs. generic drug applications)</td><td><u>Applicant Sessions</u>

Planning of application

Good Submissions for Generic Drug Applications</td></tr>
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Summary of Feedback for 2017 Workshop

- Basically, most sessions have good satisfactions.
- Suggestions for workshop organizers are summarized as follows:
  - Include **regional examples of GRM implementations** and discuss on the **challenge**
  - More training in **communications** and **critical thinking**
  - Provide more **case studies** and **interactive discussions**
  - Not only use the case of drugs, but also include **other medical products**
  - Give enough time for delegates to prepare their experience sharing

2018 APEC GRM CoE Workshop

Date :
September 26 to September 28, 2018

Agenda:
This workshop will include reviewer sessions, applicant sessions and combined sessions.

Target Audience:
Senior regulators or industry managers with at least 3 years of hands-on experiences in the management of regulatory reviews or regulatory submissions

http://www.raps-in-taiwan.org.tw/2018CoE/index.html

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Program of 2018 GRM CoE Workshop (draft)

Day 1	Day 2		Day 3
Common Sessions	***Reviewer Sessions***	***Applicant Sessions***	***Common Sessions***
Introduction of GRM **Experience sharing from different APEC member economies**	**Managing & Conducting the review** **Critical thinking and Regulatory Decision Making** *Case studies in generic drugs (1 case on CMC)*	**Planning of applications**	**Comprehensive exercises in GRM** *Preparation and submission of labeling* *Review of labeling* *Case studies and practices*
Common Sessions	***Reviewer Sessions***	***Applicant Sessions***	***Common Sessions***
Communication	**Critical thinking and Regulatory Decision Making** *Case studies in generic drugs and BE (1 case on BE)* *Case studies in biosimilar (1 case)*	**Preparation of Application Dossier/ Practice: How to Prepare Application Dossier**	**Competencies and training for reviewers and applicants** **Rolling out the GRM Training Program in each economy**

Current Focus

Step 1	Step 2	Step 3	Step 4
2011-2012	**2011-2016**	**2017-2019**	**2018-2020**
Gap Analysis Survey for setting the foundation for Stepwise GRevP Implementation	Planned solutions to address gaps GRM CoE Pilot	Assessing impact of GRM using Performance indicators	Reaching the Goal for Implementing GRM

Dissemination of GRM (continued)

- **Train-the-trainer by APEC CoE**
- **Local training by each economy**

Assessing the impact of GRM

- **Understanding the current status and challenges**
- **Developing performance indicators and assessing the impact**

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Performance Indicators for GRM

Measurable outcomes in different aspects of GRM

GRevP

- Reviewer Competency and Training
- Use of Templates and Procedures
- Transparency, Consistency, Predictability and Timeliness

GSubP

- Applicants Competency and Training
- Quality of Submission

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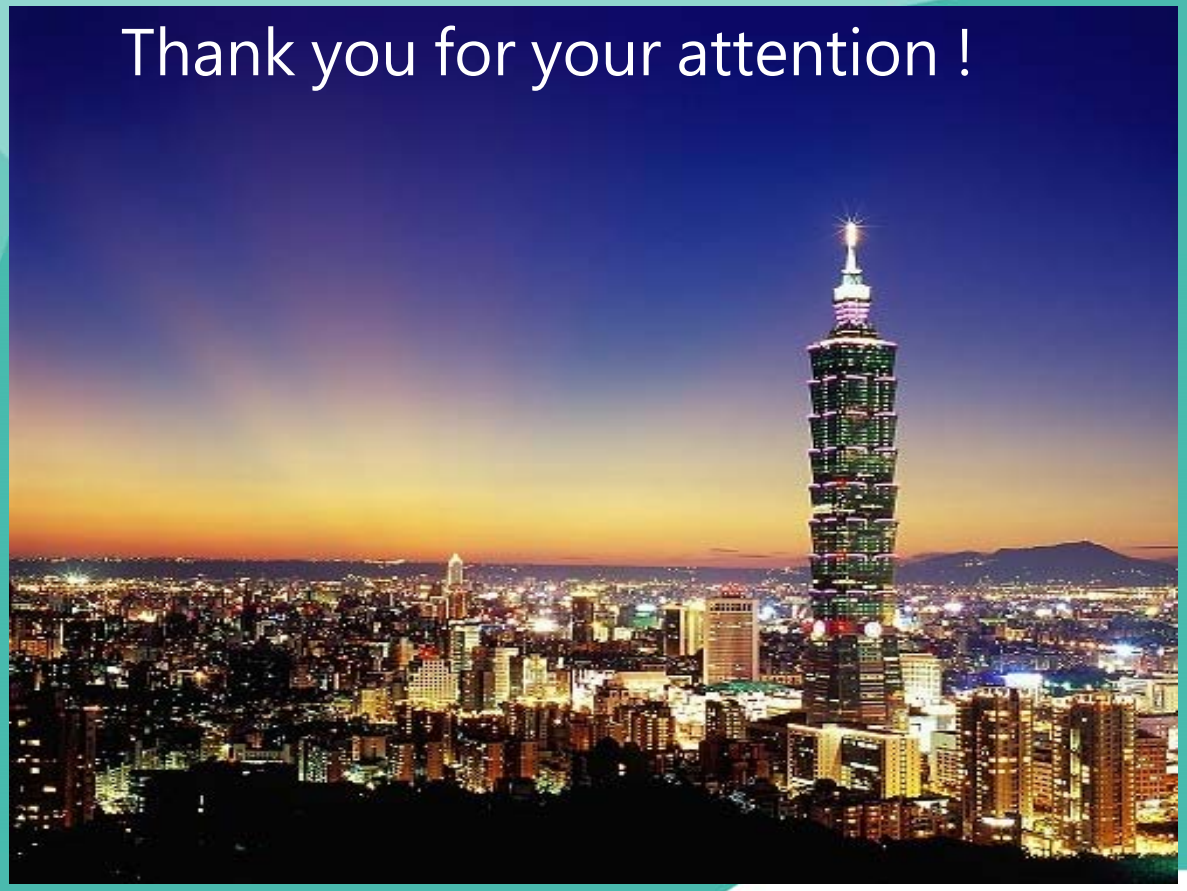
Summary (1)

- **GRM** is the concept to promote GRevP and GSubP cooperatively
- **Objective of GRM:** To enhance quality and efficiency of overall medical product registration process
- **Goal of GRM:** To benefit the patients with timely access of medicinal products of safe, efficacious and good quality
- **Guidelines for GRevP and GSubP** have been developed to provide high-level guidance on the principles and processes of good review and submission

Summary (2)

- **APEC GRM CoE** serves as a platform for promoting **regulatory convergence, capacity and cooperation** in the topic area of GRM. The focus of convergence would be in **science** and **best practices**.
- **Relevant resources** to promote GRM may include the followings:
  - Guidelines
  - Training program, materials and trainers
  - Documented best practices and quality management
  - Information sharing platform



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