APVMA Licensing For Veterinary Medicines

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Australian Government
Australian Pesticides and Veterinary Medicines Authority

To be discussed

- Commonly asked questions related to the cGMP and applicable to APVMA license holders.
- Legislation and Regulations that effect Licence Holders.
- Certificate of Export for Veterinary medicines
- Veterinary vaccines using platform technology master files (vPTMF)

What is required to gain a licence

For Aseptic manufacturers:

- Need to apply for an APVMA licence.
- Undergo an audit:
  - If TGA licensed, TGA inspection report and relevant information can be used.
  - Or an APVMA audit by an APVMA-authorised auditor to be conducted within three (3) months.

What is required to gain a licence

For APVMA Audit:

- Manufacturing facilities need to meet similar requirements as for an initial TGA inspection, with the below exception:
  - For equipment, Installation Qualification (IQ) and Operational Qualification (OQ) must be complete for critical equipment. Protocols should be available for assessment by the auditor for the future performance qualification (PQ).

APVMA legislation for GMP

Agricultural and Veterinary Chemicals Code Act 1994 (Agvet Code)

- *Contains the detailed provisions allowing the APVMA to evaluate, approve or register and review active constituents and agricultural and veterinary chemical products (and their associated labels), and to issue permits and license the manufacture of chemical products. It also contains provisions for controls to regulate the supply of chemical products, and provisions ensuring compliance with, and for the enforcement of, the Agvet Code.*

Agricultural and Veterinary Chemicals Code Regulations 1995 (Agvet Code Regs)

- These Regulations prescribe and specify a number of matters relating to the operation of the Agvet Code.

APVMA legislation for GMP

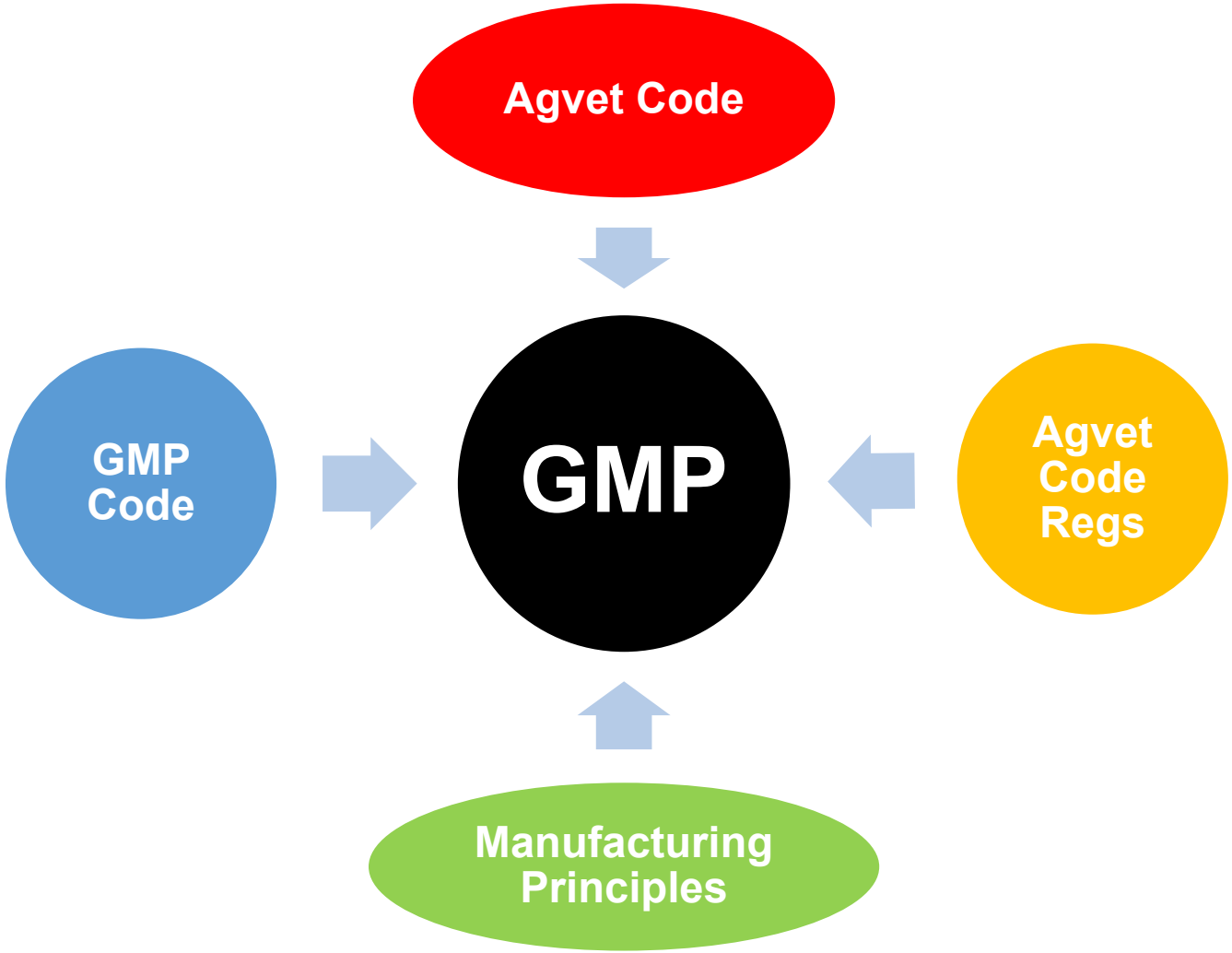
Agricultural and Veterinary Chemicals (Manufacturing Principles) Determination 2014

- This instrument sets out the principles to be observed in the manufacture of veterinary chemical products unless the products are exempt from licensing under the Agvet Regulations.

Australian code of good manufacturing practice for veterinary chemical products (Code of GMP)

- Guidance requirements to meet Good Manufacturing Practice (GMP) of veterinary chemical products.

GMP of Veterinary Chemical Products




What parts of the Act and regulations affect Licence Holders?

Licence holder responsibilities

- Under the Agvet Code Act:
- Part 8 – Manufacture of chemical products
  - Section 120 to 128
    - 120 Explanation of Part
    - 120A Exclusion of certain chemical products
    - 121 Offences relating to manufacture and licences
    - 122 Application for licence
    - 123 Issue of licence
    - 125 Period of licence
    - 126 Conditions of licences
    - 127 Suspension and cancellation of licences
    - 128 Publication of list of manufacturers etc.

Licence holder responsibilities

- Under the Agvet Code Regs:
- Part 7 – Manufacture of chemical products
  - Regulations 59 to 62
  - **59** Part 8 of the Code does not apply to listed chemical products, reserved chemical products and certain other products
  - **59A** Manufacture of chemical products—exempt persons—single step
  - **59B** Manufacture of chemical products—exempt persons—chemical product that ceases to be prescribed
  - **59C** Manufacture of chemical products—exempt persons—legal personal representative etc of licence holder
  - **59D** Manufacture of chemical products—exempt persons—person that acquires business including transfer of licence

Licence holder responsibilities

- Under the Agvet Code Regs:
- Part 7 – Manufacture of chemical products
  - Regulations 59 to 62

- **59E** Requirements for issue of licence
- **60** Licence condition—holder to give information about manufacture
- **61** Licence conditions—general
- **61A** Determination following GMP audit
- **62** Licence condition—naming persons in control of production etc

Licence holder responsibilities

- Recalls
  - Under the Agvet Code Act:
  - Part 6 – Recall notices
    - Section 100 to 107
  - Under the Agvet Code Regs:
  - Part 5A – Recall notice
    - Regulation 56A
- AERP
  - Under the Agvet Code Act:
  - Part 10 – Miscellaneous
    - Section 161
- Fees
  - Under the Agvet Code Act:
  - Part 10 – Miscellaneous
    - Section 164
  - Under the Agvet Code Regs:
  - Part 9 – Miscellaneous
    - Regulation 72A
      - Fee for licence application
      - Annual licence fee
      - Fee for variation of licence

Certificate of Export for Veterinary medicines

Agricultural and Veterinary Chemicals (Administration) Act 1992 (Admin Act 1992)

69D Export of chemical products

- These certificates are product specific, and each registered product will be issued a certificate with the current details from the APVMA register.
- Last GMP audit and GMP code.

70 Acts done by APVMA

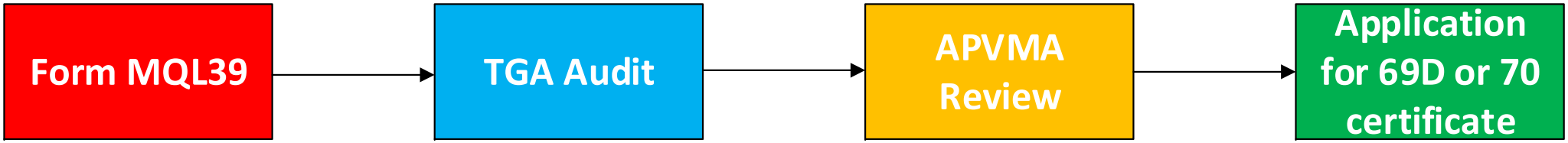
- These certificates for unregistered products, confirming that TGA have inspected these products to current PIC/S Standard of GMP.
- Last GMP audit.

PIC/S Standard of GMP

- TGA is a member of PIC/S.
- Currently 56 PIC/S participating authorities worldwide.
- Updates that will impact on veterinary manufacturers:
  - Revision of Annex 4 of the guidelines on good manufacturing practice – manufacture of veterinary medicinal products other than immunologicals.
  - Revision of annex 5 of the guidelines on good manufacturing practice for medicinal products – manufacture of immunological veterinary medicinal products.

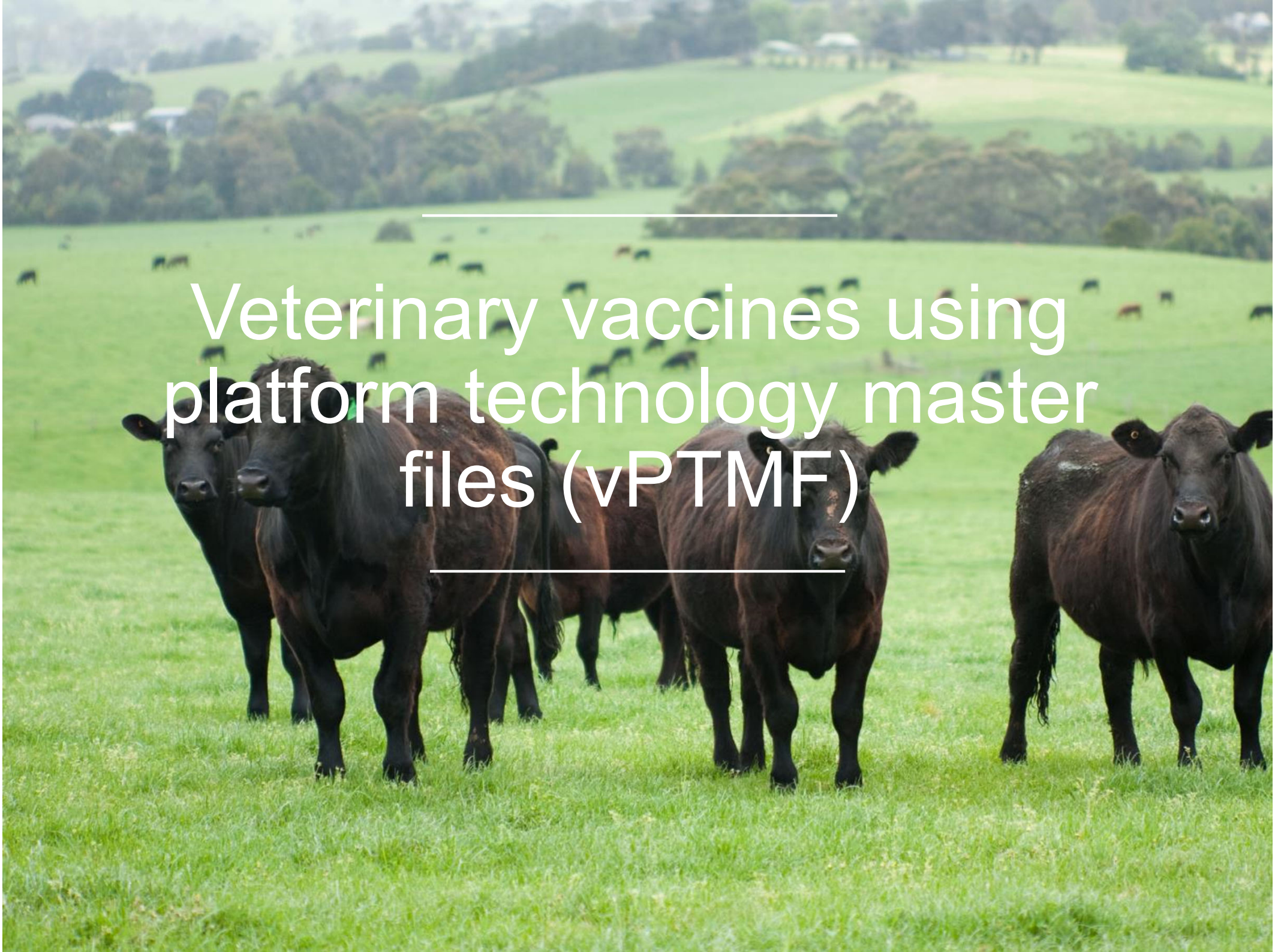
Process for TGA audit for export

- Current process has been in place since 2021.



Submit application for 69D or 70 certificate

- Applications are made through the Online Services Portal.
  - Certificate of Export and Manufacture (Approved GMP) – 69D.
  - Certificate of Manufacture (Approved GMP) – 70.
- Fee $125 for 69D applications.
- APVMA endeavours to process certificates requests within 20 working days.



Veterinary vaccines using platform technology master files (vPTMF)

Where does GMP start?

- APVMA GMP for a veterinary chemical product starts at the quality assurance of raw materials and the step of manufacture where the active material(s) and other relevant starting materials/ingredients are introduced into the formulated product and all following steps of manufacture in the final product manufacturing process.

- However, finished product to meet the s5A safety criteria of the Agvet Code and 8AB and 17C(2) of the Agvet Code Regulations.

Where Does GMP Start, Practically?

- Case by case basis depending on the active ingredient source.
- How is the active ingredient sourced?
  - Chemically synthetised or biological?
    - Chemically synthetised actives no further GMP requirements
    - Biological, further GMP controls required to be in place for safety i.e. virus and bacterial vectors and cells used are based on a seed lot system.

Where Does GMP Start, Practically?

Steps (shown in grey) used in this type of manufacturing

Biotechnology: fermentation/ cell culture	Establishment of master cell bank and working cell bank	Maintenance of working cell bank	Cell culture and/or fermentation	Isolation and purification	Physical processing, and packaging

Vaccine platform technology

- What platforms that may be used, but not limited to:
  - Protein-based platforms (virus-like particles),
  - DNA vaccine platforms,
  - mRNA-based platforms,
  - Replicons and other self-amplifying RNA; and
  - Viral and bacterial vector vaccines
- Modified cell vectors is a mature technology.
- Industry looking at more synthetic methodologies

Australian Developments

- APVMA have had several enquires / conversations with Universities.
- There are differing options on where GMP starts.
- Require an APVMA licence for steps of manufacture for permits and registration.

Key Points

- GMP for veterinary chemical products is a combination of the Agvet Code, Agvet Code Regs, Manufacturing Principles and the Code of GMP.
- TGA Inspection reports used for APVMA licence.
- Certificate of Export for Veterinary medicines.
- Veterinary vaccines using vPTMF.
- Applying or variations to licence, please contact mls@apvma.gov.au.

Thank you

Any questions?

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