#### Genetically Modified Organisms in Controlled Environments

Steve Millam SCRI, Dundee, UK "Working within the legislation"

- Context of GMs in UK
- Legislation
- Design of facilities
- Working practices
- Future

## **Plant Biotechnology**

"The advent of the techniques of plant biotechnology in the last 20 years has widened the scope and increased the precision of crop plant improvement".

**Discuss**!

#### GM plants - history

During the 1970s many attempts were made to introduce foreign DNA into plants, but all failed.

Until...

#### GM plants - history

1983, a breakthrough in the science of crop improvement published, a tobacco plant was shown to have stably integrated a gene derived from a bacteria.

> Herrera-Estrella et al., (1983) *Nature* 303, 209-213

#### GM plants - history

This was mediated using a "natural" genetic engineer, the soil bacterium *Agrobacterium*.



#### **Current Methodologies**

Other methods of gene transfer exist e.g. biolistics

Over 120 plant species have been transformed to date including potato, tobacco, oilseed rape and sugar beet.

#### **Current Methodologies**

The technology has moved very quickly since the first report.

First generation transgenics were a result of relatively unsophisticated technologies.

#### **Current Methodologies**

Recently, the methodology much more precise, and factors such as antibiotic resistance eliminated. Introduced genes may also only be switched on at specific stages of the plants life cycle.

## Historical perspective

1983 - First report published 1990 – 60+ species transformed 1992 – first field trials 1995 – first GM product on UK supermarket shelves.

## Transgenic Plant Technology

#### Year 2000

Worldwide area of GM crops reported as 34.8million ha .
16% of area for 4 crops is GM.

SOYA

COTTON





## Transgenic Plant Technology

# Strategic changes in transgenic research in the UK:

Away from the large scale commercial trials.....
Towards small scale high value or proof of principle projects.

- It is clear that the technology moves very fast.
- Scientists/Legislators have to be reactive to developments.
- Design of new facilities must have eye on future!

### The legislation

#### Covered by previous speakers

- HSE Guide to the Genetically Modified Organisms (Contained Use) Regulations 2000.
- ACGM Compendium of Guidance (Notably Parts 2D, 3B).
- Plant Health Order (1993).
- Environmental Protection Act (1990).

Notifications: HSE, & SCRI has an extra administrative layer in that Scotland is covered by SERAD.

# Containment and control measures for work with genetically modified plants

ACGM

Level A – recommended for work with plants that are unlikely to cause environmental harm.

Level B – for work where harm could arise if the GM plant (including pollen) were able to enter the environment.

# Containment and control measures for work with genetically modified plants

#### Level A – some examples

- 1. Plant is incapable of living outdoors in UK.
- 2. Plant has limited ability to transfer genetic material to UK species.
- 3. Plants transformed using a disarmed strain of *Agrobacterium* (unless conferring harmful phenotype).

# Containment and control measures for work with genetically modified plants

#### Level B – some examples

- Risk assessment identifies hazard to humans or environment e.g pollen escape.
- 2. GM plants with ability to transfer novel material to UK species.
- 3. GM plants that express hazardous substances.

#### **Containment and control measures for work with plants infected with, or in association with, GMMS**

Four levels – relevant sections Level 2:

- spatial isolation of experiments with different GMMS in association with plants employed.
- Entrance lobby with interlocking doors.
- Prevention of run-off.
- Minimisation of dispersal of pollen.
- Protective clothing.

#### **Containment and control measures for work with plants infected with, or in association with, GMMS**

Four levels – relevant sections Level 3:

- "likely to be highly engineered and expensive greenhouses".
- Sealed benches and floors.
- negative pressure.

#### Legislation – local procedures

- Genetic Modification Safety
   Committee
- Proposals submitted in advance.
- Risk assessments made.
- Committee considers each proposal and containment level.
- BSO Advises on licences etc.

 Containment measures are not only based on the use of physical barriers....

 but rely on rigorous procedural and management control.

**Physical barriers** 

- Engineering control measures.
- Supplemented with protective clothing/equipment.
- Testing and maintaining.

#### Procedural and management

- Training of personnel.
- Local codes of practice.
- Signs, notices.
- Hygiene facilities.
- Record keeping.

#### Range of crops and conditions

- Diverse face of science these days necessitates flexibility in facilities e.g.
- Arabidopsis
- Viral vectors
- GM fungi



#### Range of crops and conditions

This also indicates requirements for:

- Precise control of conditions.
- Scope for handling a number of experiments simultaneously.
- Rapid throughput and analysis.

#### Range of crops and conditions

Consequently, there is an extremely high emphasis on management, training, and detailed planning and recording of experiments.

#### SCRI -

Had been working with GMs since mid 1980s.
Working within legislation.
Facilities located at several sites across Institute.

### SCRI -

- Plans for centralising facilities for GM research, (or building a custom-designed facility) had been around since early 1990s.
- However, a major funding opportunity arose, which with the assistance of SET, University of Dundee, ERDF.

### SCRI -

- Came up with £1.6 million for a purpose built lab, growth room and containment glasshouse facility.
- Diverse sources of funding necessitated that teaching and business aspects featured.

## SCRI – planning

- Consultation with scientists as to future needs (!).
- Legislative factors strongly featured.
- Visits to other sites/related facilities (GM containment learns from phytosanitary set-ups).

## SCRI – planning

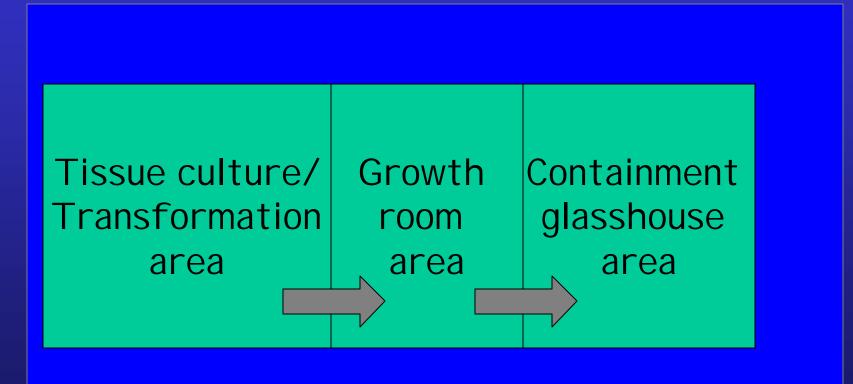
Finalised a flexible package: -

- Primary focus on protecting plants & protecting environment.
- Facility designed to deal with experiments ranging from single plant to pre-field scale.

## SCRI – planning

- Efficient ergonomic flow of work.
- Computer control & recording.
- Range of environments within.
- Distinctive.
- All within a "limited" budget!!

- SERAD consultation at early plan stages to ensure features for current and future legislation were incorporated.
- Run through worked examples of current GM projects.



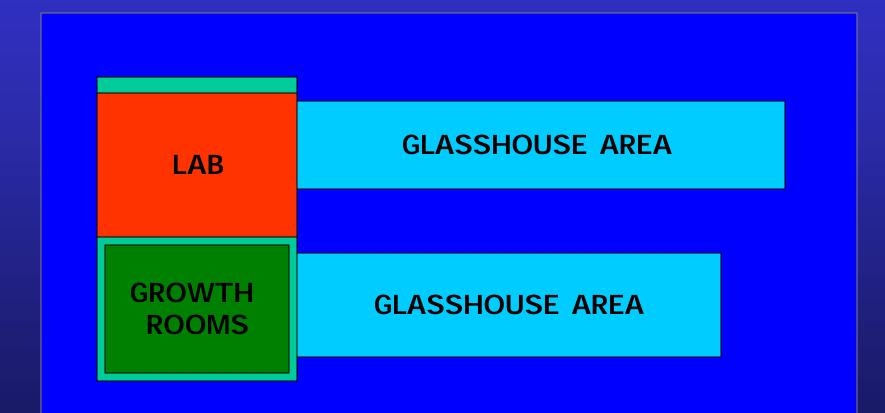


#### Lab/Plant room



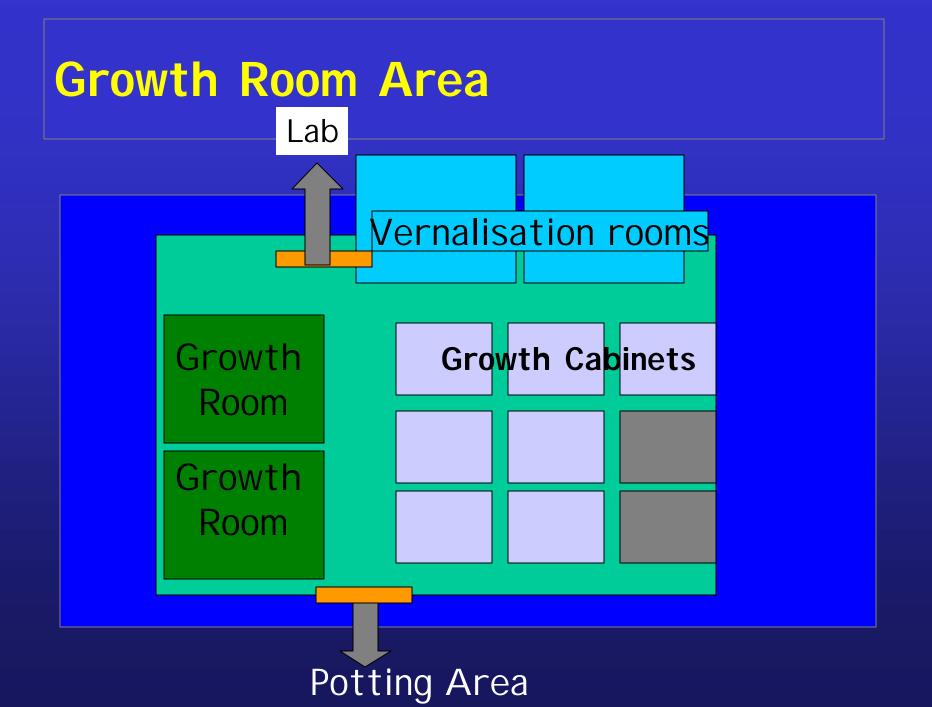
#### South aspect





# Working within the legislation: *lab area*

Access limited to key holders only
 Logical throughput of work
 Agrobacterium area isolated
 Designated Growth rooms
 Waste material disposal
 Lab records



## Working within the legislation: growth room area

- Access to key holders only
- All cabinets/rooms lockable
- All have experimental details (inc. contacts, licence number)
- Waste disposal procedures
- P&D programme

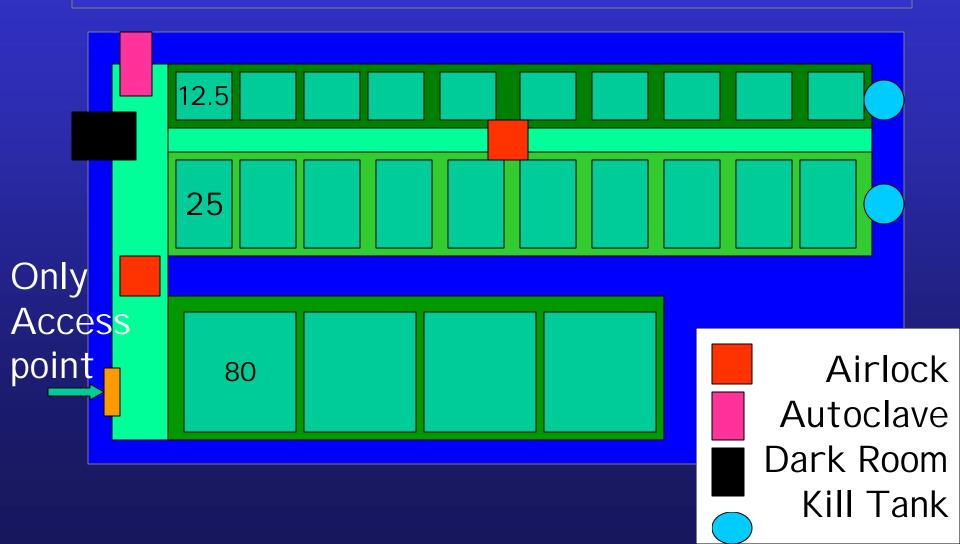
### Working within the legislation: growth rooms





Range of species grown include *Arabidopsis*, Potato, *Ribes* and barley

#### **Glasshouse** Area



## Working within the legislation: glasshouse area

- All drains sealed/sealable
- Appropriate filters on fan units
- All compartments fully sealed
- All compartments lockable
- Autoclaves in each designated area
- Protective clothing/hygiene
- SOPs

## Working within the legislation: glasshouse area





#### low containment area

Autoclave in medium containment area



## Working within the legislation: glasshouse area



Air lock between high and medium areas



Sealed containers for transport "colour-coded" trolleys

### Working within the legislation: glasshouse area - features

Changes since initial design

- Designated Glasshouse Staff
- Enhanced S.O. Procedures
- Wider range of projects
- Dark room facility added

#### Working within the legislation: Future prospects.....

#### • ISO9000

- Increase in viral vector work
- Increased range of species
- Special Arabidopsis provision
- High throughput systems
- In situ analytical systems

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#### Acknowledgements:



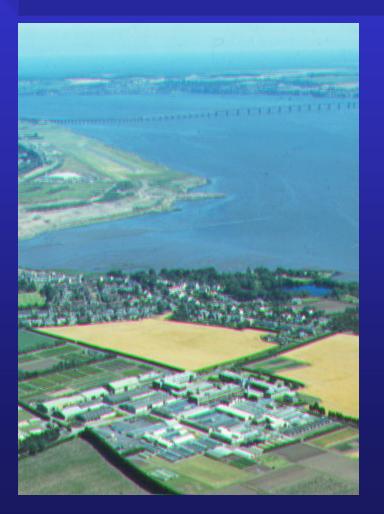


ERDF/SET/ Dundee & Abertay Universities.

SCRI is funded by SEERAD

#### **Further information**





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