

RB264

on
Bew 92 A

BSE - TRANSMISSION STUDIES

DATES OF INOCULATION AND RESULT/ONSET OF CLINICAL SIGNS
CHARACTERISTIC OF A SCRAPIE-LIKE DISEASE

IA BSE COW BRAIN - To various host species

| | | <u>Place</u> | <u>Route</u> | <u>Exposure</u> | <u>Status</u> | <u>Rounded Minimum Incubation Period</u> |
|-----------|-----|------------------------|--------------|--|-------------------------------------|---|
| MICE | NPU | I/C I/P | | Nov/Dec 1987 | Clin. Sep 88 | 9.7M (292d) |
| MICE | RVC | Oral | | Early 1988 | Clin. 1989 | 15M |
| CATTLE | CVL | I/C I/P | | Jan/May 1988 | Clin. Jul 89 | 18M (8.6M*) |
| HAMSTERS | CVL | I/C I/P | | Feb/May 1988 | All dead -ve ^{oo} | No distinction between challenge and controls |
| MARMOSETS | CRC | I/C | | Feb 1988 | Healthy | |
| PIGS | CVL | I/C I/P I/V Oral | | 24 Feb/5 Mar 1989 25 May/7, 15 Jun 1990 | Clin. Jun 90 Healthy | 16M (481d) |
| SHEEP | NPU | I/C Oral | | 24 Jun 1988 12 Aug 1988 | Clin. Aug 89 Please see sheet IB | 14M |
| GOATS | NPU | I/C Oral | | 24 Jun 1988 4 Aug 1988 | Clin. Dec 89 Healthy | 18M |
| CHICKEN | CVL | I/C I/P Oral (Crop) | | 5/18 Jun 1990 6, 11, 18 Jul 1990 | Healthy Healthy | |
| MINK | USA | I/C Oral | | 21 Feb 1991 21 Feb 1991 | Healthy Healthy | |

Clin. clinical positive confirmed by histopathology

* onset of prodromal phase

oo in confidence (unpublished)

PLACENTA

CATTLE CVL Oronasal Nov 1989 Healthy

As at 30 April 1991

In Confidence

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BSE TRANSMISSION EXPERIMENTS - PROGRESS TO DATE (30.4.91)

Source Material - Bovine Brain

| IB | <u>PARENTERAL</u> | A <u>CONTROLS</u> | B <u>CHALLENGE</u> | <u>REMARKS</u> |
|----|-------------------|---------------------------------------|---|--|
| | Cattle | 8 alive | 16 dead | A healthy B positive BSE |
| | Sheep | | a) 2/6 dead - line b) 2/5 dead + line | a) 1 positive scrapie 1 negative pathology b) 1 negative pathology 1 pending |
| | Goats | | 3 dead 0 alive | B positive scrapie |
| | Chickens | 13 alive 1 dead | 9 alive 3 dead | A 1 dead SAF -ve (sudden death) rest healthy B 3 dead SAF -ve rest healthy |
| | Pigs | 5 alive 2 dead 4 killed @ 2Y | 4 alive 2 dead 1 clinical 3 killed @ 2Y | A 2 intercurrent deaths 4 killed pending B 2 intercurrent deaths 1 confirmed positive 3 killed pending |
| | Marmosets | | 2 alive (BSE) | B healthy |
| | Mink | Inoculated | 21/2/91 USA | A, B healthy |

| IC | <u>ORAL</u> | A <u>CONTROLS</u> | B <u>CHALLENGE</u> | <u>REMARKS</u> |
|----|-------------|----------------------|--|--|
| | Sheep | | a) 1/6 dead + line b) 1/8 dead - line | a) 1 positive scrapie (? natural) b) clinical negative positive pathology |
| | Goats | | | B All healthy |
| | Chickens | As above | 11 alive 3 dead | B 3 died at dosing rest healthy |
| | Pigs | 10 alive | 10 alive | A, B all healthy |

Source Material - Bovine Placenta

| ID | <u>ORAL</u> | A <u>CONTROLS</u> | B <u>CHALLENGE</u> | <u>REMARKS</u> |
|----|-------------|----------------------|-----------------------|------------------|
| | Cattle | 9 alive | 12 alive | A, B all healthy |

II INFECTIVITY OF TISSUES FROM CATTLE WITH BSE OR ANIMALS WITH
FOR MICE

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Route of Inoculation I/C 0.02 ml I/P 0.1 ml 10% suspension

| | <u>Place</u> | <u>Exposure</u> | | <u>Results/</u> <u>Clinical +ve</u> | <u>Minimum</u> <u>Incubation</u> <u>Period</u> |
|-----------------------|--------------|-----------------|-----------------|--|--|
| SEmen | NPU | 3 Aug | 1989 (Tox) | | |
| | - | 30 Aug | 1989 | | |
| SPLEEN | NPU | 3 Aug | 1989 | | |
| | | 4 Sept | 1989 | | |
| BUFFY COAT | NPU | 1 Nov | 1989 | | |
| M DIAPHRAGMA | NPU | 9 Nov | 1989 | | |
| M SEMITENDINOSUS | NPU | 6 Nov | 1989 | | |
| COW BRAIN X 2 | NPU | 2 Nov | 1989 | | |
| | | 7 Nov | 1989 | | |
| FIXED COW BRAIN X 2 | NPU | 8 Nov | 1989 | | |
| PLACENTA | NPU | 2 Feb | 1990 | | |
| MILK (TOXICITY TRIAL) | | | | | |
| MILK | | | | | |
| MIDRUM | | | | | |
| LIVER | NPU | 11 Jan | 1991 | | |
| UDDER | | | | | |
| MES. LYMPH NODE (LN) | NPU | 9 Nov | 1990 | | |
| CSF | NPU | 10 Jan | 1991 | | |
| KIDNEY | | | | | |
| *EMBRYOS | | | | | |
| *UTERINE FLUSHINGS | | | | | |
| SEmen (2nd SAMPLE) | | | | | |
| PANCREAS | NPU | 24 Apr | 1991 | | |
| DISTAL ILEUM | | | | | |
| PREFEMORAL LN | NPU | 9 Nov | 1990 | | |
| PROXIMAL COLON | NPU | 24 Apr | 1991 (1/50 i/p) | | |
| RUMEN (PILLAR) | NPU | 15 Feb | 1991 | | |
| FETAL CALF BLOOD | NPU | 29 Nov | 1991 | | |
| BONE MARROW (STERNUM) | NPU | 5 Mar | 1990 | | |

* not available in sufficient quantity yet

As at 28 April 1991

III INFECTIVITY OF TISSUES FROM BSE ANIMALS FOR MICE BY ORAL EXPOSURE

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| <u>Place</u> | <u>Exposure</u> | <u>Results/</u> | | <u>Minimum Incubation Period</u> |
|------------------|-----------------|---------------------|-------------|--|
| | | <u>Clinical +ve</u> | <u>1989</u> | |
| BRAIN + CSF | RVC | Early 1988 | 1989 | 15M |
| UDDER + MILK | RVC | Early 1988 |) | |
| SUPRAMAMMARY LN | RVC | Early 1988 |) | |
| PLACENTA | RVC | Early 1988 |) | |
| SPLEEN | RVC | Early 1988 |) | |
| CARCASE + MES LN | RVC | Early 1988 |) * | |
| | | |) | |
| | | |) | |
| SPLEEN | NPU | 27 Jan 1990 |) | |
| PLACENTA x 2 | NPU | 2 Feb 1990 |) | |

* (1) All negative

(2) 12 month parenteral passage using :-

- a) spinal cord
- b) spleen

All negative

(3) 18 month parenteral passage as in (2)

All negative at 30/4/91 but incomplete

As at 30 April 1991

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IV INFECTIVITY OF BRAIN FROM SPECIES OTHER THAN CATTLE
FOR MICE OR OTHER SPECIES

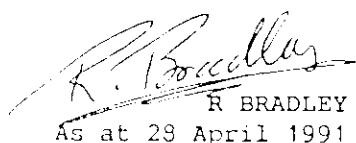
| <u>Source</u> | <u>Place</u> | <u>Route</u> | <u>Exposure</u> | <u>Status</u> | <u>Min i/p</u> |
|-------------------|--------------|--------------|-----------------|---------------|----------------|
| FIXED NYALA BRAIN | NPU | i/c i/p | 27 Dec 1989 | Positive | 343d. |
| FIXED KUDU BRAIN | NPU | i/c i/p | 28 Dec 1989 | Positive | 302d. |
| CATS | NPU | - i/c i/p | 22 Aug 1990 | Healthy | |
| | | | 26 Sep 1990 | " | |
| | | | 16/29 Jan 1991 | " | |
| CWD | NPU | i/c i/p | 5 Mar 1991 | " | |
| EXPTL PIG SE | NPU | i/c i/p | 30 Oct 1990 | " | |

SCRAPIE

| | | | | |
|-------------------|-----|---------|------------|----------------------------|
| GREY FACE TO MICE | NPU | i/c i/p | 8 Jan 1988 | Positive (longer than BSE) |
| TO MARMOSETS | CRC | i/c | Feb 1988 | Positive 38M |

OTHER POSITIVE EXPERIMENTS

| | | | | |
|-----------------------|-----|-------------|-------------|----------|
| FORMALIN/BSE - MICE | NPU | i/c i/p | 8 Nov 1989 | Positive |
| | | | 9 Nov 1989 | Positive |
| 1M NaOH 263 K TO MICE | NPU | i/c | 29 Sep 1990 | Positive |
| 2M NaOH 263 K TO MICE | NPU | i/c | 16 Sep 1990 | Positive |
| RIII BSE TO HAMSTER | NPU | i/c or | | |
| | | i/p or both | 4 Apr 1990 | Positive |
| VM BSE TO HAMSTER | NPU | " | 3 Apr 1990 | Positive |
| -ve LINE CHEVIOT BSE | | | | |
| - MICE | NPU | i/c i/p | 27 Apr 1990 | Positive |



R. Bradley
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As at 29 April 1991

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