Table 2-1 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs: home cage observations (Week 1)

| | Sex | | Male | | | | Female | | | |
|---------------------------|----------------|----|------|-----|------|----|--------|-----|------|--|
| | Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 | |
| Parameter | No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Posture Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Convulsion None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Abnormal behavior None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |

| • | Sex | | Male | | | Female | | | | |
|---------------------------|----------------|----|------|-----|------|--------|-----|-----|------|--|
| | Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 | |
| Parameter | No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Posture Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Convulsion None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Abnormal behavior None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |

| | Sex | | Male | | | | Female | | | |
|---------------------------|----------------|----|------|-----|------|----|--------|-----|------|--|
| | Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 | |
| Parameter | No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Posture Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Convulsion None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Abnormal behavior None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |

Table 2-4 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs: home cage observations (Week 4)

| | Sex | | Ma | ıle | | Female | | | |
|---------------------------|----------------|----|-----|-----|------|--------|-----|-----|------|
| | Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 |
| Parameter | No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Posture Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Convulsion None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Abnormal behavior None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |

| | Sex | Male | | Female | | |
|---------------------------|----------------|------|------|--------|------|---|
| | Dose (mg/kg) | 0 | 1000 | 0 | 1000 | _ |
| Parameter | No. of animals | 6 | 6 | 6 | 6 | |
| Posture Normal | | 6 | 6 | 6 | 6 | |
| Convulsion None | | 6 | 6 | 6 | 6 | |
| Abnormal behavior None | | 6 | 6 | 6 | 6 | |

| | Sex | Male | | Female | | |
|---------------------------|----------------|------|------|--------|------|---|
| | Dose (mg/kg) | 0 | 1000 | 0 | 1000 | • |
| Parameter | No. of animals | 6 | 6 | 6 | 6 | |
| Posture Normal | | 6 | 6 | 6 | . 6 | • |
| Convulsion None | | 6 | 6 | 6 | 6 | |
| Abnormal behavior None | | 6 | 6 | 6 | 6 | |

Table 2-7 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs: in-the-hand observations (Week 1)

| Sex | | Ma | ıle | | | Fen | nale | |
|---|--------------|-------------|-------------|--------------|--------------|-------------|-------------|--------------|
| Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 |
| Parameter No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Ease of removal from cage | 1.1 | 0 | e | 10 | 10 | 6 | 0 | 1.1 |
| Easy Some resistance/avoidance Difficult | 11 1 0 | 6 0 0 | 6 0 0 | 10 1 1 | 12 0 0 | 5 1 0 | 6 0 0 | 11 1 0 |
| Fur condition Normal | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Skin Normal | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Secretions-Eye, Nose Absent | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Exophthalmos Absent | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Palpebral closure Normal | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Mucosal membranes Normal | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Lacrimation Normal | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Piloerection Absent | 12 | 6 | 6 | 12 | 12 | 6 | . 6 | 12 |
| Pupil size Normal | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Salivation None | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Abnormal respiration Absent | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 |
| Reactivity to handling Easy Slightly awkward Difficult | 10 1 1 | 6 0 0 | 6 0 0 | 10 2 0 | 12 0 0 | 5 1 0 | 6 0 0 | 11 1 0 |

Table 2-8 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs: in-the-hand observations (Week 2)

| | | | | | | | | | | |
|-------------------------------|----------------|-------------|-----|-----|------|----|------------|-----|------|---|
| | Sex | | Ma | le | | | Fem | ale | | |
| | Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 | |
| Parameter | No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Ease of removal fro | om cage | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Fur condition Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Skin Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Secretions-Eye, Nos Absent | se | 12 | 6 | 6 | 12 | 12 | 6 , | 6 | 12 | |
| Exophthalmos Absent | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Palpebral closure Normal | | 12 | 6 | 6 | 12 . | 12 | 6 | 6 | 12 | |
| Mucosal membranes Normal | | 12 | 6 | 6 | 12 | 12 | 6 - | 6 | 12 | • |
| Lacrimation Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Piloerection Absent | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Pupil size Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Salivation None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Abnormal respiration | nc | 12 | 6 | 6 | 12 | 12 | . 6 | 6 | 12 | |
| Reactivity to hand Easy | ling | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |

Table 2-9 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs: in-the-hand observations (Week 3)

| | Sex | | Ma | ıle | | | Fem | ale | | |
|--|----------------|---------|-----|-----|---------|---------|--------|--------|---------|--|
| | Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 | |
| Parameter | No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Ease of removal Easy Some resistance | - | 12 0 | 6 | 6 | 12 0 | 11 1 | 6 0 | 6 0 | 12 0 | |
| Fur condition Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Skin Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Secretions-Eye, Absent | Nose | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Exophthalmos Absent | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Palpebral closum Normal | ·е | 12 | . 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Aucosal membrane Normal | es | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Lacrimation Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Piloerection Absent | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Pupil size Normal | | 12 | 6 | в | 12 | 12 | 6 | 6 | 12 | |
| Salivation None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Abnormal respira Absent | ation | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Reactivity to ha | andling | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |

Table 2-10 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs: in-the-hand observations (Week 4)

| | Sex | | Ma | le. | | | Fem | ale | | |
|--------------------------------|----------------|----|-----|-----|------|----|-----|-----|------|--|
| | Dose (mg/kg) | 0 | 100 | 300 | 1000 | 0 | 100 | 300 | 1000 | |
| Parameter | No. of animals | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Ease of removal from | cage | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Fur condition Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Skin Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Secretions-Eye, Nose Absent | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Exophthalmos . Absent | | 12 | 6 | 6 | 12 | 12 | 6 · | 6 | 12 | |
| Palpebral closure Normal | | 12 | 6 | 6 | .12 | 12 | 6 | 6 | 12 | |
| Mucosal membranes Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Lacrimation Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Piloerection Absent | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Pupil size Normal | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Salivation None | | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Abnormal respiration Absent | ı | 12 | 6 | 6 | 12 | 12 | 6 | 6 | 12 | |
| Reactivity to handli Easy | ng | 12 | 6 | 6 , | 12 | 12 | 6 | 6 | 12 | |

Table 2-11 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs: in-the-hand observations (Week 1 of recovery)

| | Sex | M | ale | Fe | emale |
|--------------------------------------|----------------|--------|--------|--------|--------|
| | Dose (mg/kg) | 0 | 1000 | 0 | 1000 |
| Parameter | No. of animals | 6 | 6 | 6 | 6 |
| Ease of removal | from cage | 6 | 6 | 6 | 6 |
| Fur condition Normal | | 6 | 6 | 6 | 6 |
| Skin Normal | | 6 | 6 | 6 | 6 |
| Secretions-Eye, Absent | Nose | 6 | 6 | 6 | 6 |
| Exophthalmos Absent | | 6 | 6 | 6 | 6 |
| Palpebral closur Normal | re | 6 | 6 | 6 | 6 |
| Mucosal membrane Normal | es | 6 | 6 | 6 | - 6 |
| Lacrimation Normal | | 6 | 6 | 6 | 6 |
| Piloerection Absent | | 6 | 6 | 6 | 6 |
| Pupil size Normal | | 6 | 6 | 6 | 6 |
| Salivation None | | 6 | 6 | 6 | 6 |
| Abnormal respira Absent | ation | 6 | 6 | 6 | 6 |
| Reactivity to he Easy Slightly awkwa | | 6 0 | 6 0 | 5 1 | 6 0 |

Table 2-12 A 28-day oral toxicity study of paracetaldehyde in rats with a recovery period of 2 weeks

Detailed clinical signs : in-the-hand observations (Week 2 of recovery)

| | Sex | М | ale | Fe | male | | |
|----------------------------|----------------|---|------|-----|------|------|------|
| • | Dose (mg/kg) | 0 | 1000 | . 0 | 1000 | | |
| arameter | No. of animals | 6 | 6 | 6 | 6 | | |
| ase of removal f Easy | rom cage | 6 | 6 | 6 | 6 | | |
| ır condition Normal | | 6 | 6 | 6 | 6 | | |
| in Normal | | 6 | 6 | 6 | 6 | | |
| ecretions-Eye, N Absent | ose | 6 | 6 | 6 | 6 | | |
| ophthalmos Absent | | 6 | 6 | 6 | 6 | | |
| alpebral closure Normal | | 6 | 6 | 6 | 6 | | |
| ıcosal membranes Normal | | 6 | 6 | 6 | 6 | | |
| crimation Normal | | 6 | 6 | 6 | 6 | | |
| loerection Absent | | 6 | 6 | в | 6 | | |
| pil size Normal | | 6 | 6 | б | 6 | | |
| alivation None | | 6 | 6 | 6 | 6 | | |
| normal respirat Absent | ion | 6 | 6 | 6 | 6 | | |
| activity to han Easy | dling | 6 | 6 (| 6 | 6 | | |