Disruption of Brain Development and Reproductive Behavior of Birds

DM Fry, JR Millam, AL Erichsen, MA Viant, AE Quaglino, CB Craig-Veit, and LA Preston. Dept of Animal/Avian Sci. University of Calif. Davis

Extrapolation of xenobiotic exposure to the "real world"

- Methoxychlor, dicofol, and alkylphenols in agriculture in Yolo Co.
- Application rates and acreages available.
- Sensitivity of local species unknown



Current research on songbirds

- Altricial chicks require extended parental care
- Complex, but well understood brain neuroanatomy and physiology
- Song system is estrogen sensitive
- Brain is dimorphic



Zebra Finches

Hormonal Influence in Brain Differentiation

- Testosterone produced by neo-natal male gonads
- Uptake and aromatization by specific areas of brain
- Estrogen modulates differentiation of male specific nuclei. Mechanism uncertain, BDNF clearly involved, apoptosis?

Dosing and testing protocol

- Oral gavage
- 5-11 days post hatch
- 1µ1 / g body wt. / day in canola oil
- parental care of chicks for 30-40 days
- mixed sex juvenile cages

- Behavioral testing at 120+ days
- Behavior assessed with stimulus males and females on alternate days
- Testosterone implants to force/enable altered behavior
- Histology of brain, gonad

Effective doses given to finches (µg/g/body wt./day)

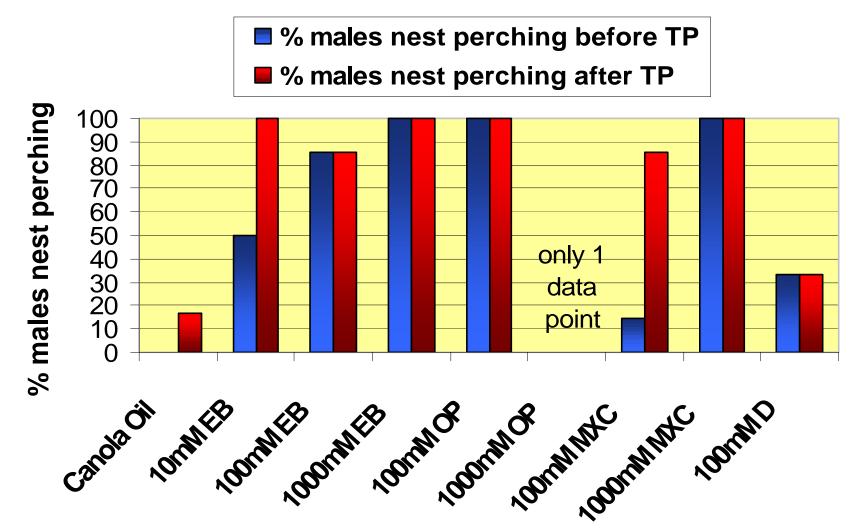
- Estradiol benzoate: 2.7
- Methoxychlor: 35
- Octylphenol: 21
- Dicofol: 37



Extrapolated dose to nestling songbirds

- Body wt. During sensitive period: 3-10g
- Food consumption: 1-8 g insects / day
- Dicofol @ 189 ppm,
 Exposure dose is 189-1512µg/day
- Octylphenolpolyehoxylate @ 22ppm: Exposure dose is 22-176µg/day

"Broodiness" of Male Zebra Finches



Summary of Effects on Males

- Estradiol and Xenoestrogens active orally
- Microgram per day doses cause changes in male default behavior patterns:
 - Latency for courtship song
 - Reduced copulation
 - "Broodiness"
 - Responses in both

Zebra Finches and Japanese Quail



Differentiation of brain and behavior in birds:

Song System:

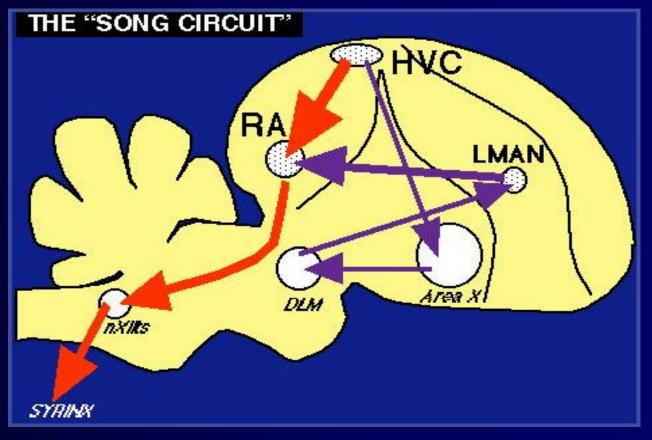
- Passerine birds only
- Species specific patterns in dimorphism
- Estrogen sensitive during post-natal development



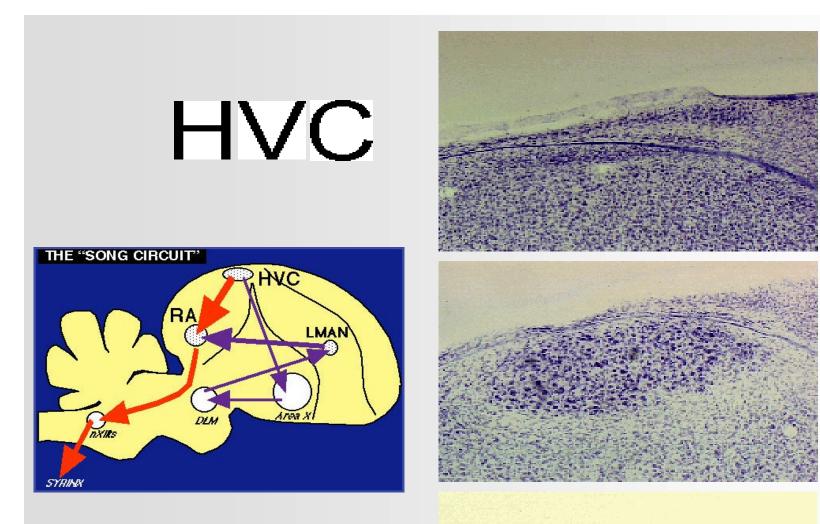
Female Zebra Finch

Zebra Finch "Song Circuitry"

- Learning and memory pathways in blue
- Motor pathway in red
- Males sing, females do not



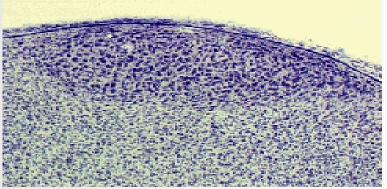
David Clayton, Univ Illinois



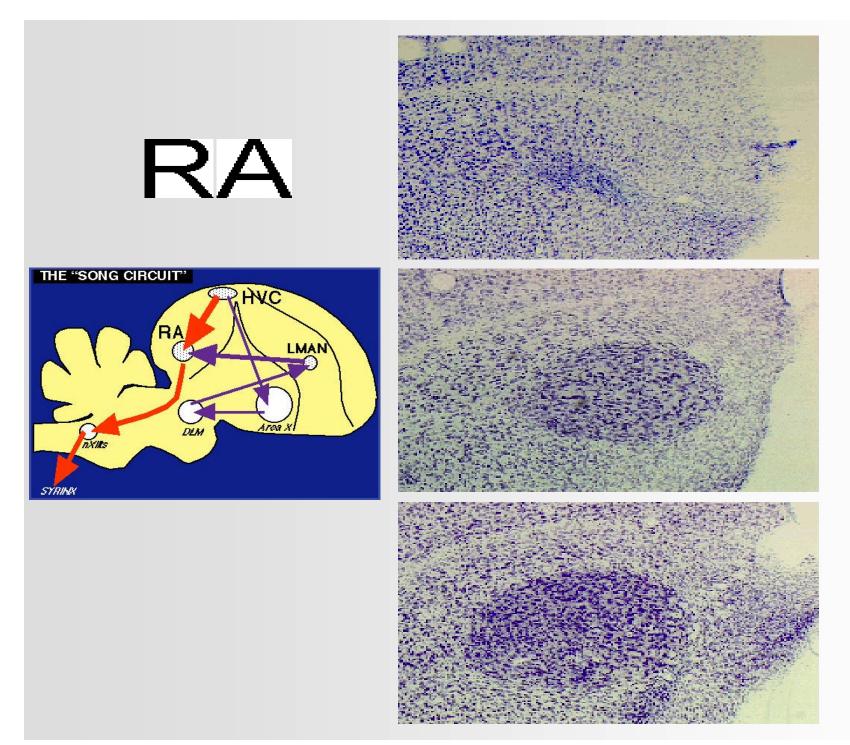
Canola Oil Female

100mM EB Female

Threshold for enabling song is about $125\mu m^3$



Canola Oil Male



Canola Oil Female

100mM EB Female

Canola Oil Male

Vocalizations of captive Zebra finches

Male Zebra Finch

M Distance Calls

Chatter ("tet" call)

Courtship song

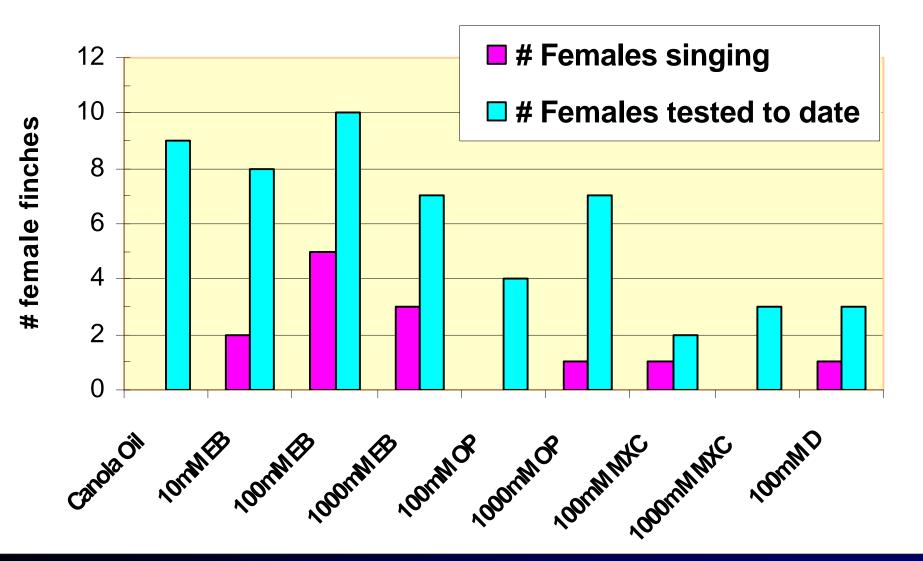


Vocalizations of female zebra finches

- Normal Females
 "tet" call
- 🍕 distance call
- Male type" songs bysensitive exposed females



Female Finches Singing Courtship Song



Estrogen and xenoestrogen induced changes in female finches

- Sexually dimorphic brain nuclei induced (retained? / enhanced?)
- Singing enabled in exposed females
- Aggressive behavior in many females
- Significant phenotypic variability by family



Reproductive Performance Tests

- Neonatal Dosing of zebra Finches, days 5-11
 - Estradiol benzoate 27µg/gBW/day
 - Octylphenol 21µ g/gBW/day
- Group caging: 6 pair per cage
- Dosed males and females together
- Dosed birds with control mates
- Reproductive activity from Day 140

Reproductive Activity:

- Estradiol females lay eggs, clutch size similar
- No eggs hatch from treated group
 - dosed males and females together
 - eggshell thinning in most females
- Mixed pair trials in progress:
 - some fertility and hatching success.