# CANCER AND LIFE HISTORY THEORY

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#### CAVEATS AND CONSTRAINTS

- I am an PhD Evolutionary Psychologist, not an MD Physician:
  - And I don't even play one on TV!
- Given those constraints, I will attempt to derive general principles from Life History Theory that might specifically relate to the other talks in this session on the subject of *Cooperation, Conflict,* and Co-evolution:
  - Drawing explicit analogies from psychosocial phenomena addressed in my own program of research on Human Life History Strategy

- Williams, Miller, Harper, & Wiersma (2010) have provided both comparative and experimental evidence in birds that the rate of *cell metabolism* is directly related to the rate of *cell division*
- Controlling for body weight, tropical bird species have lower metabolic rates than do temperate bird species
- Controlling for body weight, tropical bird species have longer life spans than do temperate bird species

- Cell-cultured Dermal fibroblasts derived from tropical birds have slower rates of growth than do such cells from birds of temperate species:
  - Consistent with the hypothesis that these cells have a slower metabolism
- Conversely, cell-cultured dermal fibroblasts derived from tropical birds resist chemical agents that induce oxidative and non-oxidative stress better than do such cells from birds of temperate species:
  - Consistent with the hypothesis that birds that live longer invest more in self-maintenance, such as in the antioxidant properties of cells

- Joyce and Pollard (2009) have provided experimental evidence that the microenvironment in which tumors develop play a major role in modulating the metastatic capacity of most cancers:
  - The formation of metastases has many ratelimiting steps, and certain *microenvironmental cues* are important moderating influences at each and every one of these rate-limiting steps

- The implication is that the different microenvironments within the body in which tumors develop may systematically select for different allocations of bioenergetic and material resources towards:
  - Faster metabolism, growth, and proliferation
  - Self-maintenance, resistance, and survival

- Elser, Kyle, Smith, & Nagy (2007) proposed the Growth Rate Hypothesis (GRH), in which a tumor growing in the body can be considered a complex ecological and evolutionary system
- The protein synthesis demands of accelerated cell proliferation in tumors produce elevated phosphorus demands, due to increased allocations to phosphorus-rich nucleic acids such as ribosomal RNA
- This suggests the preferential but differential allocation of resources in tumor tissues towards the numerical proliferation as opposed to the individual survival of cancer cells

- This bioenergetic tradeoff is analogous to those respectively characterizing r-selection (favoring allocations towards propagule proliferation) and K-selection (favoring allocations towards propagule survival)
- This theoretical prediction was tested by examining the different allocations made by cancer cells developing in different parts of the body
- The biochemical compositions of cancer tissues were examined and compared in the Liver, Kidney, Colon, and Lung

- Unstable local conditions in the Lung and Colon impose high levels of external mortality on tumor cells, producing long-term selection favoring transformed (r-selected) cells with increased rates of cell division:
  - These tumor tissues are *more* enriched in phosphorus and nucleic acids, as predicted by the *GRH*
- Stable local conditions in the Liver and Kidney may instead predominantly favor transformed (Kselected) cells with lower rates of cell mortality, such as a reduction in apoptosis:
  - These tumor tissues are *less* enriched in phosphorus and nucleic acids, as predicted by the *GRH*

### SO WHAT ELSE CAN WE LEARN FROM LIFE HISTORY THEORY?

- The conditions for the evolution and development of fast and slow life history strategies have been extensively studied:
  - Reviewed in Ellis, Figueredo, Brumbach, & Schlomer (2009)
- Many of these ecological conditions have strong implications, both directly and indirectly, for the dynamics of *Cooperation*, *Conflict*, and Co-evolution:
  - So let's review the basic mechanisms...

### LIFE HISTORY STRATEGY IS ABOUT RESOURCE ALLOCATION

- Somatic Effort:
  - Bioenergetic and material resources devoted to continued survival of individual organism
- Reproductive Effort:
  - Bioenergetic and material resources devoted to production of new organisms as vehicles for survival of individual's genes

#### REPRODUCTIVE EFFORT

- Mating Effort:
  - Bioenergetic and material resources devoted to obtaining and retaining sexual partners
- Parental/Nepotistic Effort:
  - Bioenergetic and material resources devoted to enhancing survival of any offspring produced by self or kin

# FAST AND SLOW LIFE HISTORY STRATEGIES

- Fast ("r-Selected") Life History Strategies allocate resources preferentially to:
  - Reproductive Effort over Somatic Effort
  - Mating Effort over Parental/Nepotistic Effort
  - Emphasize the *production* of new propagules over the *survival* of existing ones (whether self or offspring)

# FAST AND SLOW LIFE HISTORY STRATEGIES

- Slow ("K-Selected") Life History Strategies allocate resources preferentially to:
  - Somatic Effort over Reproductive Effort
  - Parental/Nepotistic Effort over Mating Effort
  - Emphasize the survival of propagules (whether self or offspring) over the production of new ones

# SLOW ("K-SELECTED") LIFE HISTORY STRATEGY

- As compared to Fast Life History Strategists, Slow Life History Strategists should therefore:
  - Manifest benefits of allocations in greater phenotypic quality and survivorship of propagules (whether cells or organisms)
  - Manifest greater individual viability on indicators of general health, developmental stability, and behavioral function

### RABBITS AND ELEPHANTS: DIFFERENT LIFE HISTORIES

- Rabbits (Fast Life History Strategy):
  - Rapid sexual development
  - High fertility
  - Little parental care per offspring
  - High infant mortality
  - Adults are relatively short-lived



#### Female Rabbits:

"abandon their young in burrows immediately after birth and return to feed them for only about two minutes daily during their first 25 days. After this brief bout of 'drive-by' parenting, young rabbits are left to fend for themselves."

#### RABBITS AND ELEPHANTS: DIFFERENT LIFE HISTORIES

- Elephants (Slow Life History Strategy):
  - Very slow and delayed sexual development
  - Produce few babies at a time
  - High parental care per offspring
  - Very low infant mortality
  - Adults are very long-lived



#### Female elephants:

"are pregnant for 21 months. they feed their child milk for up to 6 yrs"

"stay with their birth family for life while males live with the group until reaching puberty, between the age of nine and fifteen."

# THE EVOLUTION OF LIFE HISTORY STRATEGIES

- Fast Life History Strategies are naturally selected in unstable, unpredictable environments:
  - Sources of mortality predominantly extrinsic, and hence uncontrollable by geneticallyinfluenced developmental processes
  - Leading to highly variable population densities and reinforcing this selective effect

# THE EVOLUTION OF LIFE HISTORY STRATEGIES

- Slow Life History Strategies are naturally selected in stable, predictable environments:
  - Sources of mortality predominantly intrinsic, and hence controllable by geneticallyinfluenced (and hence evolvable) developmental processes
  - Leading to highly stable population densities and reinforcing this selective effect

### MULTIPLE LEVELS OF SELECTION FOR LIFE HISTORY

- A Hierarchical Cascade of Consequences:
  - Natural Selective Pressures generate both Individual and Social Sequelae
  - Producing Social Selective Pressures that generate Sexual Sequelae
  - Producing Sexual Selective Pressures that generate further Sexual Sequelae

### SOCIAL SELECTION FOR FAST LIFE HISTORY STRATEGIES

- Unstable, unpredictable, and uncontrollable social relationships, under which individuals (both self and others) tend to evolve and develop:
  - Insecure attachment to kith and kin
  - Opportunistic and exploitative interpersonal styles
  - Low kin-selected altruism
  - Low parental and nepotistic effort
  - High social defection
  - High social antagonism
  - High social aggression
  - Generally selfish orientation to social partners

### SOCIAL SELECTION FOR FAST LIFE HISTORY STRATEGIES

- These Socially Selected Sequelae are jointly due to:
  - The severely limited time horizon available for social, nepotistic, and parental relations due to the adverse natural selective pressures specified
  - The self-reinforcing nature of these effects (as in the Pianka, 1970, theory of *r*- and *K*-selection) due to the adverse social selective pressures in environments where the majority of conspecifics are also pursuing fast life history and short-term social strategies

### SOCIAL SELECTION FOR SLOW LIFE HISTORY STRATEGIES

- Stable, predictable, and controllable social relationships, under which individuals (both self and others) tend to evolve and develop:
  - Secure attachment to kith and kin
  - Mutually and reciprocally rewarding interpersonal styles
  - High kin-selected altruism
  - High parental and nepotistic effort
  - High social reciprocity
  - High social mutualism
  - Generally altruistic orientation to social partners

### SOCIAL SELECTION FOR SLOW LIFE HISTORY STRATEGIES

- These Socially Selected Sequelae are jointly due to:
  - The more distant and foreseeable time horizon available for social, nepotistic, and parental relations due to the relatively safe and favorable natural selective pressures specified
  - The self-reinforcing nature of these effects (as in the Pianka, 1970, theory of r- and K-selection) due to the relatively safe and favorable social selective pressures in environments where the majority of conspecifics are also pursuing slow life history and long-term social strategies

### SEXUAL SELECTION FOR FAST LIFE HISTORY STRATEGIES

- Unstable, unpredictable, and uncontrollable sexual relationships, under which individuals (both self and others) tend to evolve and develop:
  - Insecure attachment to sexual partners
  - Higher mating effort in the service of multiple short-term pairings, whether simultaneous or serial or both
  - Opportunistic and exploitative sexual relations, including deceptive and coercive sexuality
  - High cross-sexual defection
  - High cross-sexual antagonism, including low cross-sexual cooperation and coparenting
  - High cross-sexual aggression, including "intimate partner violence"
  - Generally selfish orientation to sexual partners

### SEXUAL SELECTION FOR FAST LIFE HISTORY STRATEGIES

- These Sexually Selected Sequelae are jointly due to:
  - The severely limited time horizon available for sexual relationships due to the adverse natural and social selective pressures specified
  - The self-reinforcing nature of these effects (as in the Pianka, 1970, theory of r- and K-selection) due to the adverse social and sexual selective pressures in environments where the majority of conspecifics are also pursuing fast life history and short-term social and sexual strategies

### SEXUAL SELECTION FOR SLOW LIFE HISTORY STRATEGIES

- Stable, predictable, and controllable sexual relationships, under which individuals (both self and others) tend to evolve and develop:
  - Secure attachment to romantic partners, not just sexual partners
  - Lower mating effort in the service of a reduced number of long-term pairings, perhaps not perfectly monogamous
  - Mutually and reciprocally rewarding sexual relations
  - Strong and committed cross-sexual bonds
  - High cross-sexual reciprocity
  - High cross-sexual mutualism, including cross-sexual cooperation and coparenting
  - Low cross-sexual antagonism
  - Generally altruistic orientation to sexual partners

### SEXUAL SELECTION FOR SLOW LIFE HISTORY STRATEGIES

- These Sexually Selected Sequelae are jointly due to:
  - The more distant and foreseeable time horizon available for sexual relationships due to the relatively safe and favorable natural and social selective pressures specified
  - The self-reinforcing nature of these effects (as in the Pianka, 1970, theory of r- and k-selection) due to the safe and favorable social and sexual selective pressures in environments where the majority of conspecifics are also pursuing slow life history and short-term social and sexual strategies

### SO WHAT CAN CANCER RESEARCH LEARN FROM LIFE HISTORY THEORY?

- The ecological conditions for the evolution and development of fast and slow life history strategies have strong implications for the dynamics of Cooperation, Conflict, and Coevolution:
  - The generative *Natural* Selective Pressures constrain and shape the *Social* Selective Pressures, which, in turn, constrain and shape the *Sexual* Selective Pressures that drive both Evolution and Development

### SO WHAT CAN CANCER RESEARCH LEARN FROM LIFE HISTORY THEORY?

- The implications of all these theoretical considerations for the evolution and development of cancer tissues have yet to be worked out
- The implications for the prevention or treatment of cancer is an even more distant objective

### SO WHAT CAN CANCER RESEARCH LEARN FROM LIFE HISTORY THEORY?

- Nevertheless, we may now reasonably ask:
  - What is a malignant tumor, other than a fast life history (r-selected) cancer tissue?
  - What is a benign tumor, other than a slow life history (K-selected) cancer tissue?
- If we can understand the microenvironmental cues triggering the developmental switches involved in a tumor evolving or developing into a malignant or benign one, that may be half the battle!