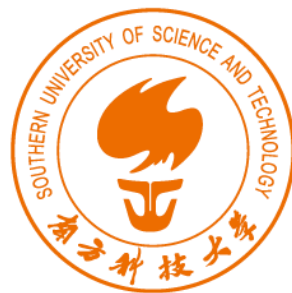


# A novel biophysical model for radical pair mechanism in birds' magnetoreception



2019年5月21日

骆锦威 11410163



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# Background

# First discovery in magnetoreception

## **Magnets Interfere with Pigeon Homing**

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*Communicated by Donald R. Griffin, October 8, 1970*

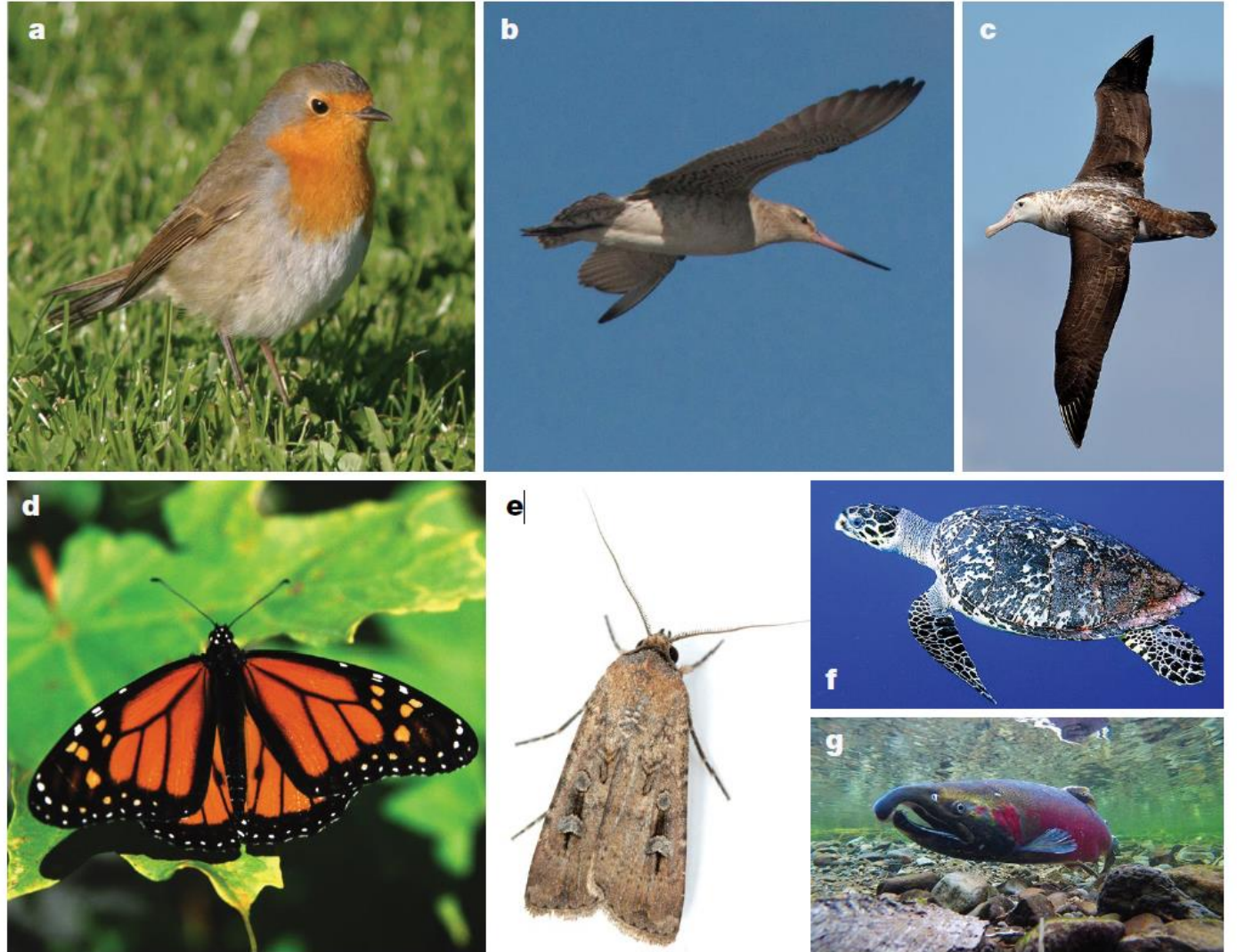
**ABSTRACT** Magnets glued to the backs of experienced pigeons often resulted in disorientation when the birds were released from distances of 17-31 miles (27-50 km) under total overcast, whereas no such disorientation occurred during similar releases under clear skies. The magnets did, however, often cause disorientation when first-flight birds were released under sun, and there was some indication of disturbance to experienced pigeons released under sun at longer distances.

# Magnetic compass is ubiquitous across species

**Vertebrates:** birds, fishes, amphibians, reptiles, mammals.

**Invertebrates:** mollusks, crustaceans, insects.

- *Erithacus rubecula*
- *Limosa lapponica*
- *Diomedea exulans*
- *Danaus plexippus*
- *Agrotis infusa*
- *Eretmochelys imbricate*
- *Oncorhynchus kisutch*



Modified after Ref.[2]



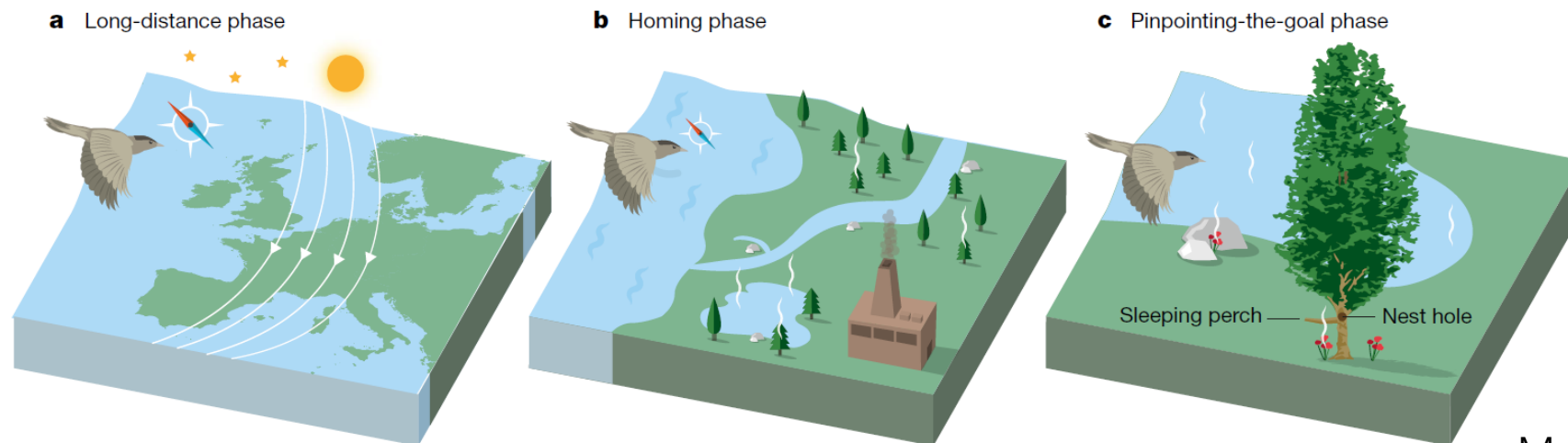
# Multisensory cues for bird navigation

- **Long-distance navigation – Migration maps**

Magnetic compass, sun compass, star compass

- **Homing and pinpointing – Local orientation**

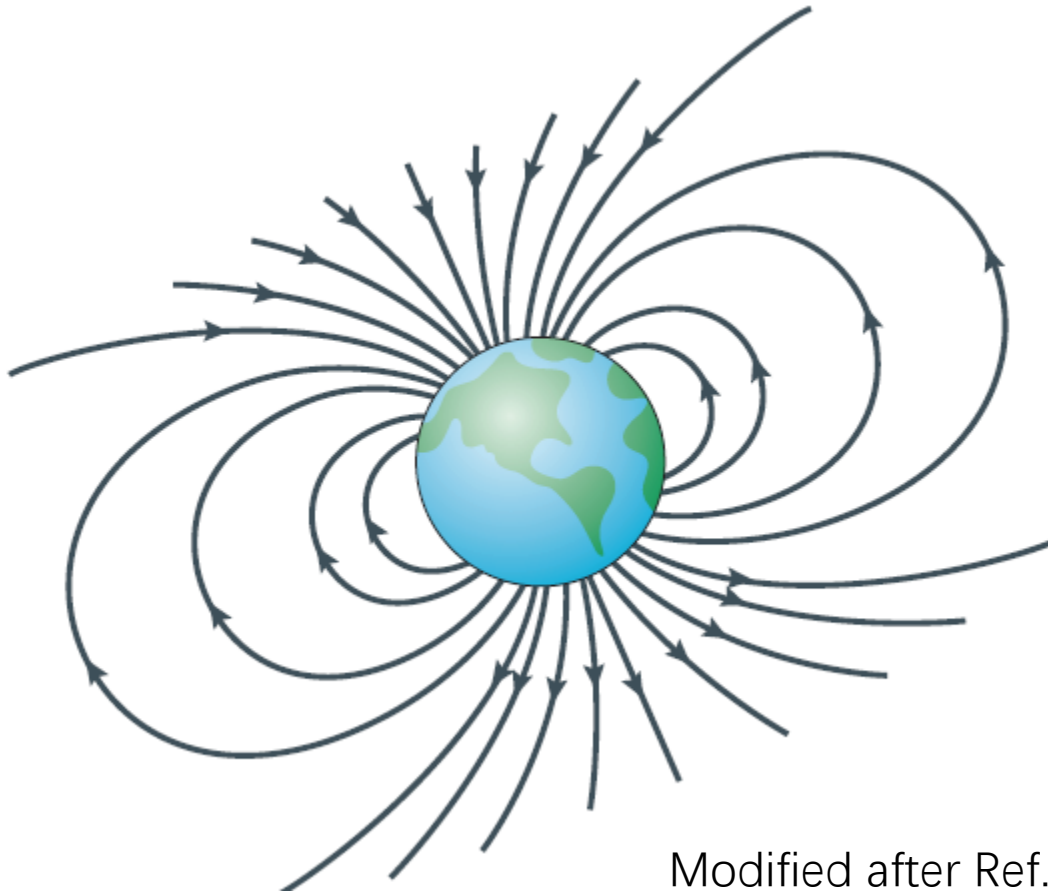
Odors, borders, coast lines, sound, landmarks such as a tree, a small hill, a specific coral



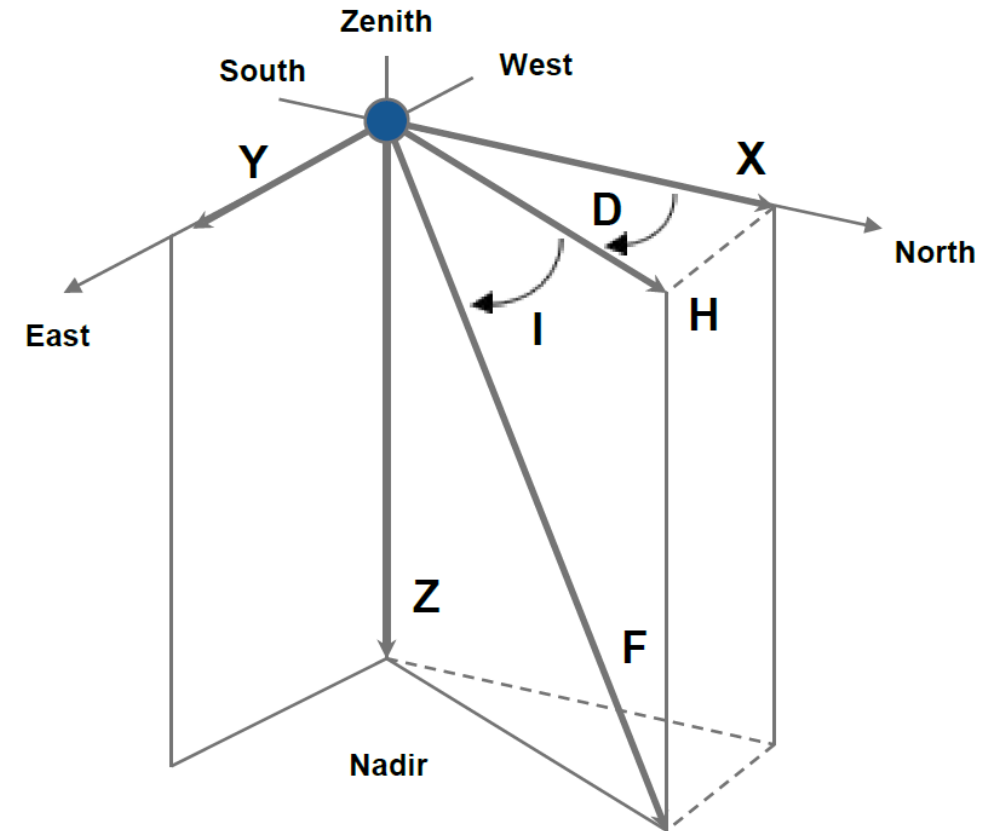
Modified after Ref.[2]

# Earth magnetic fields

- Field strength: 25~60uT



Modified after Ref.[3]



Magnetic field information

**Inclination**

**Polarity/Declination**

**Intensity**

# Mechanisms

## Three possible mechanisms

- Electromagnetic induction
- Magnetite based magnetoreception
- Radical based magnetoreception

## Difficulties for verification

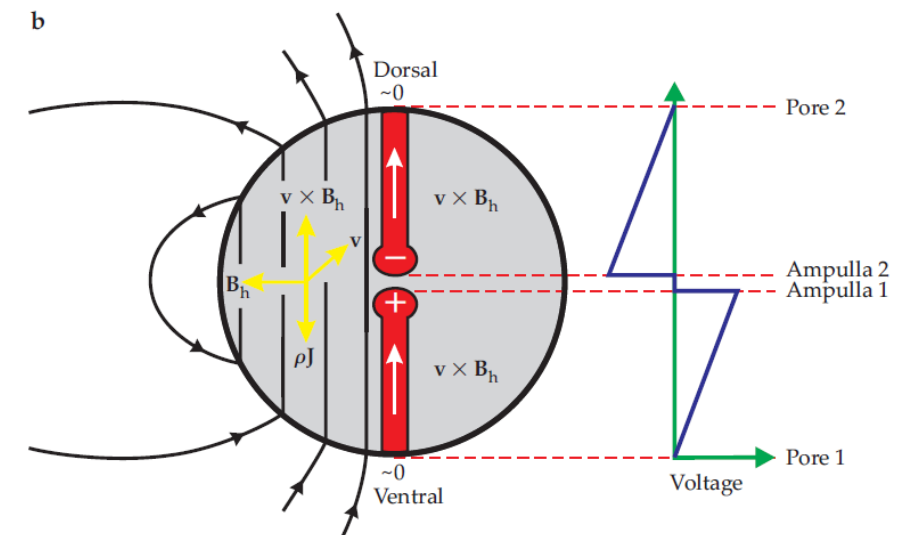
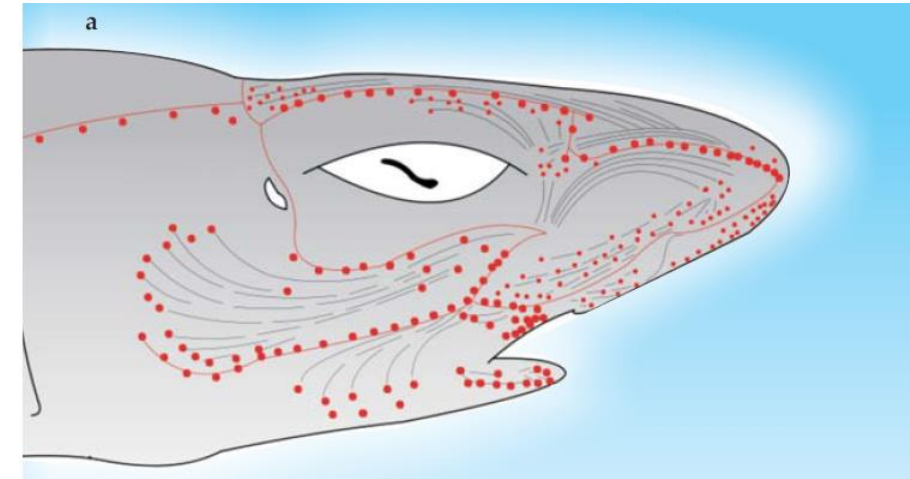
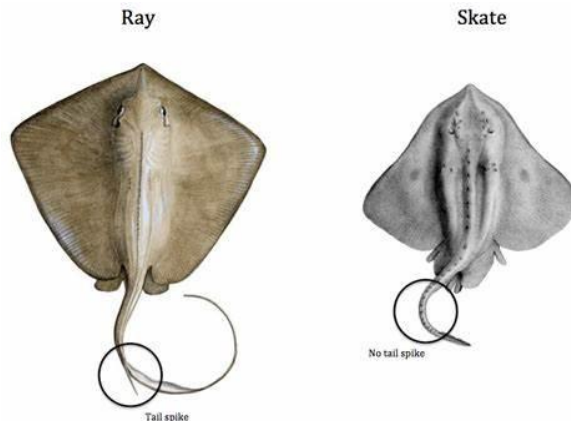
- No specific organ is found
- No unique behavioral phenotype
- Multisensory cues coupling
- Adaptation and acclimation
- Multi origins or single origin?



# Electromagnetic induction

Magnetic information -> Electric signal

- A unique organ in cartilaginous fish:
- **Ampullae of Lorenzini**
- Jelly-filled pores with electroreceptors
- Moving magnetic field accumulates electric potential on two ampullas via proton flux, which can be detected by electroreceptor cells.
- Elasmobranch species: **sharks, rays and skates**.

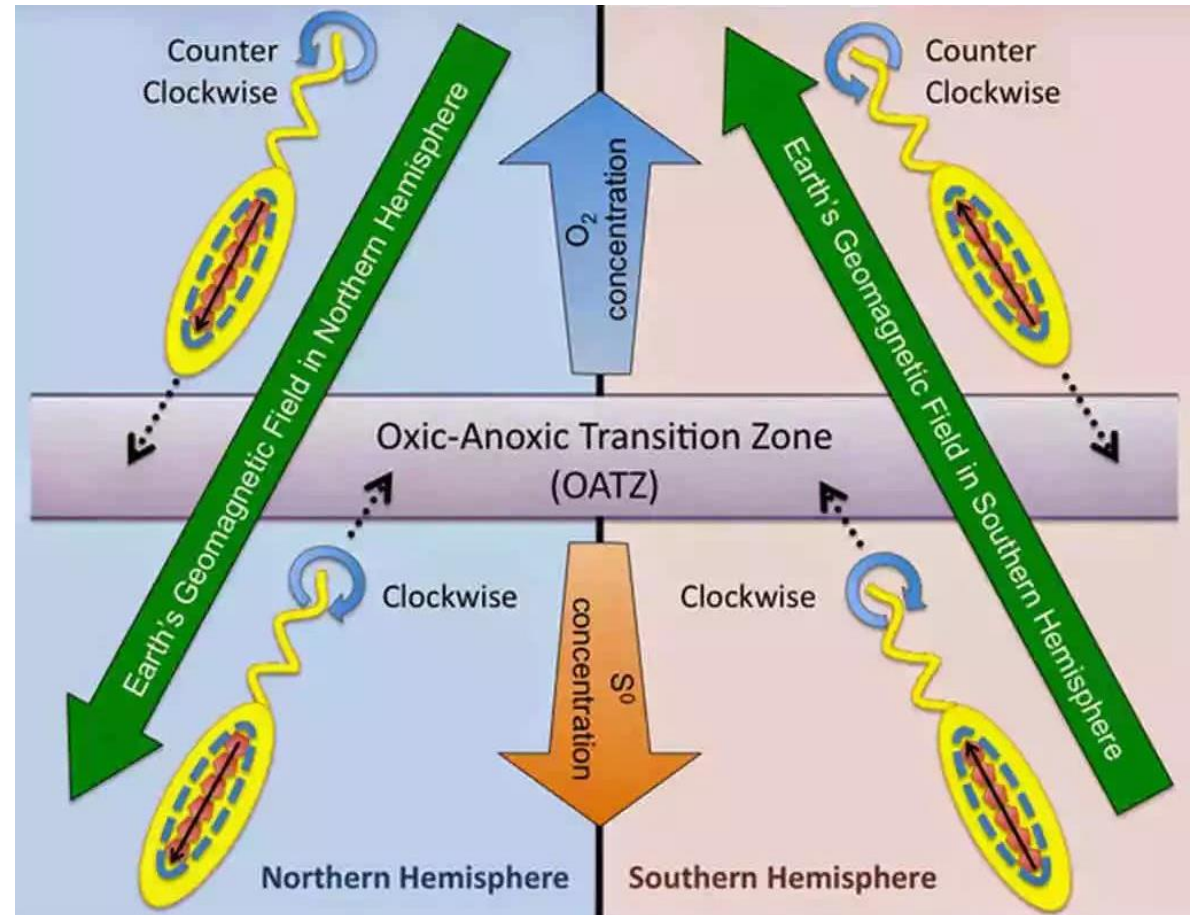
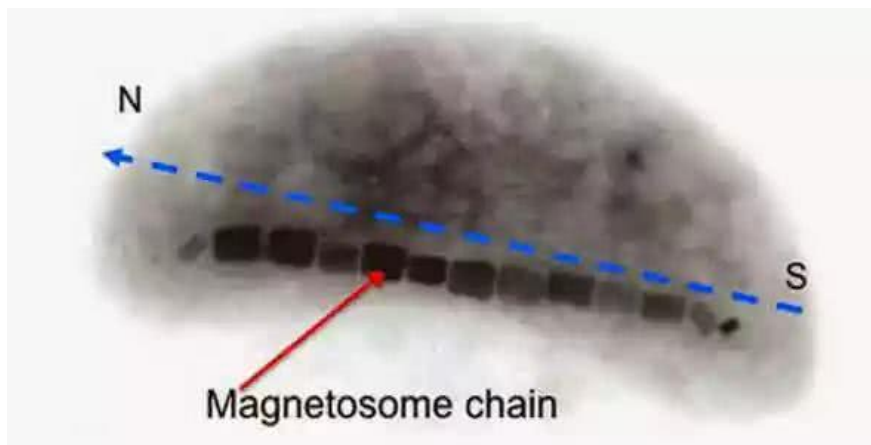


Modified after Ref.[4]

# Magnetite based Magnetoreception

Magnetic force -> Physical rotation

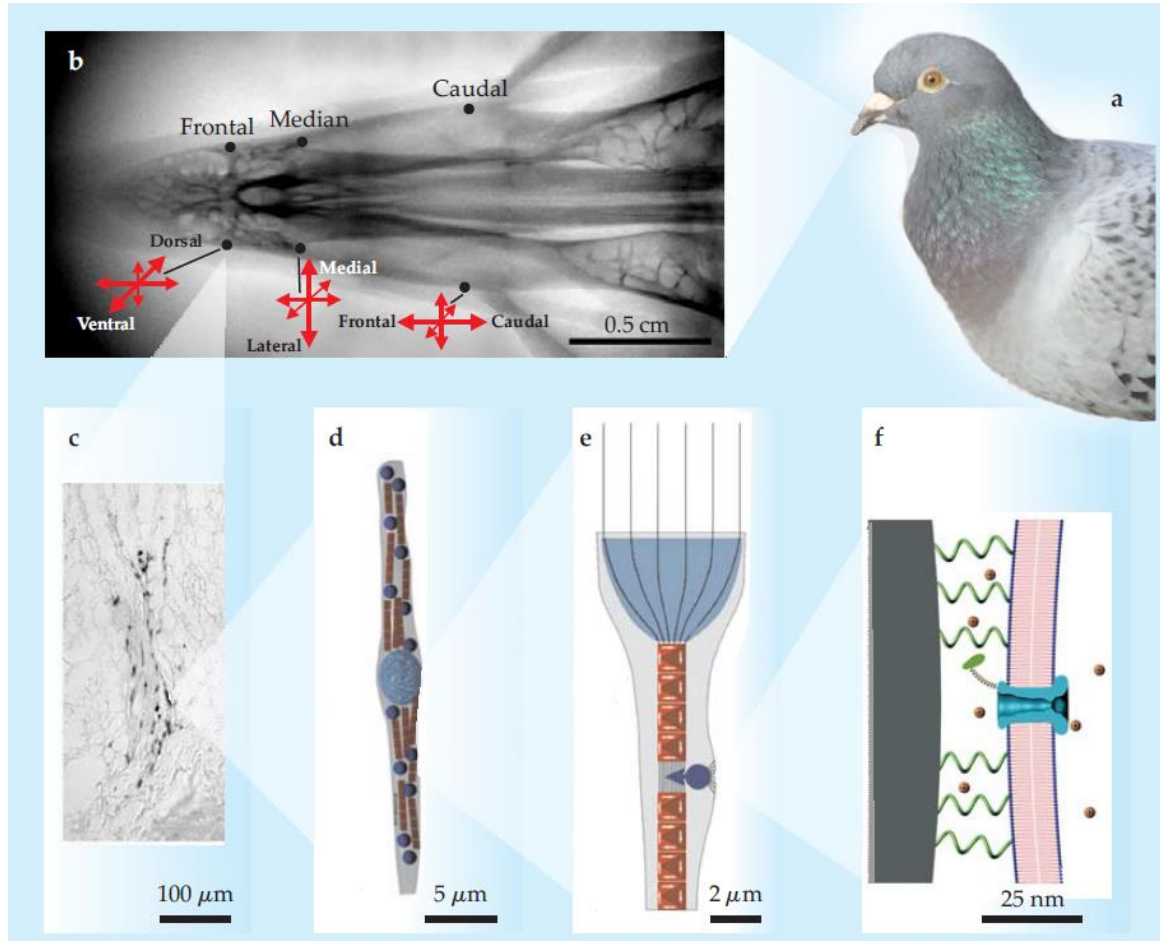
- Magnetotactic bacteria
- A magnetosome chain:
- 15-20  $\text{Fe}_3\text{S}_4$ ,  $\text{Fe}_3\text{O}_4$  crystal
- 30-100nm for each crystal



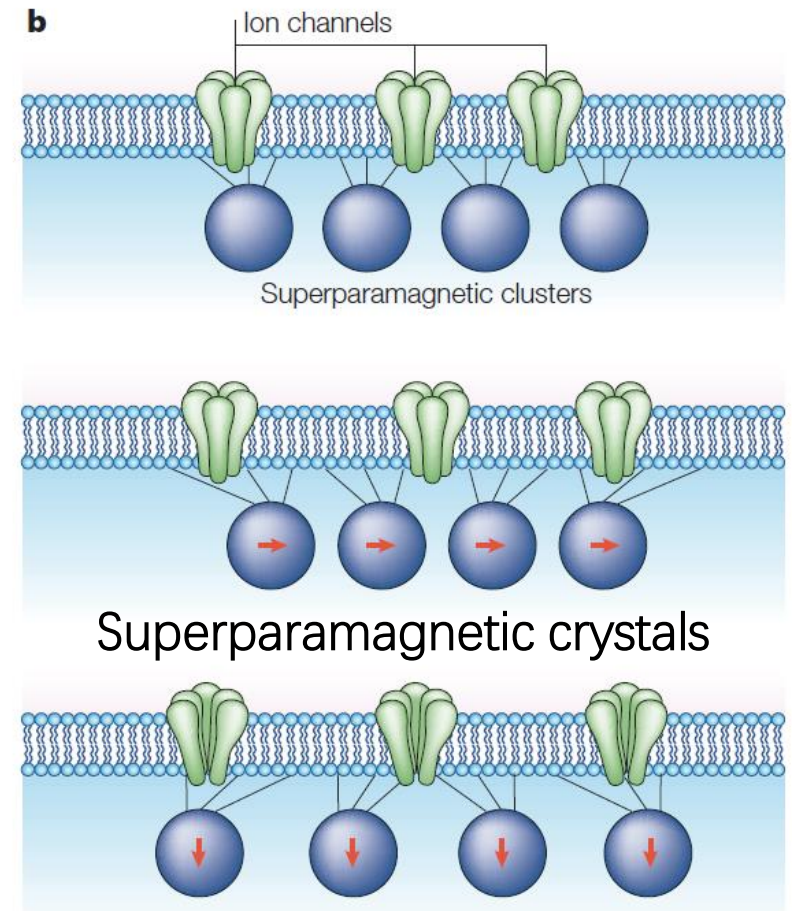
Modified after Ref.[5]

# Magnetite based Magnetoreception

Magnetic information -> Pressure signal -> Electric signal



Modified after Ref.[4]



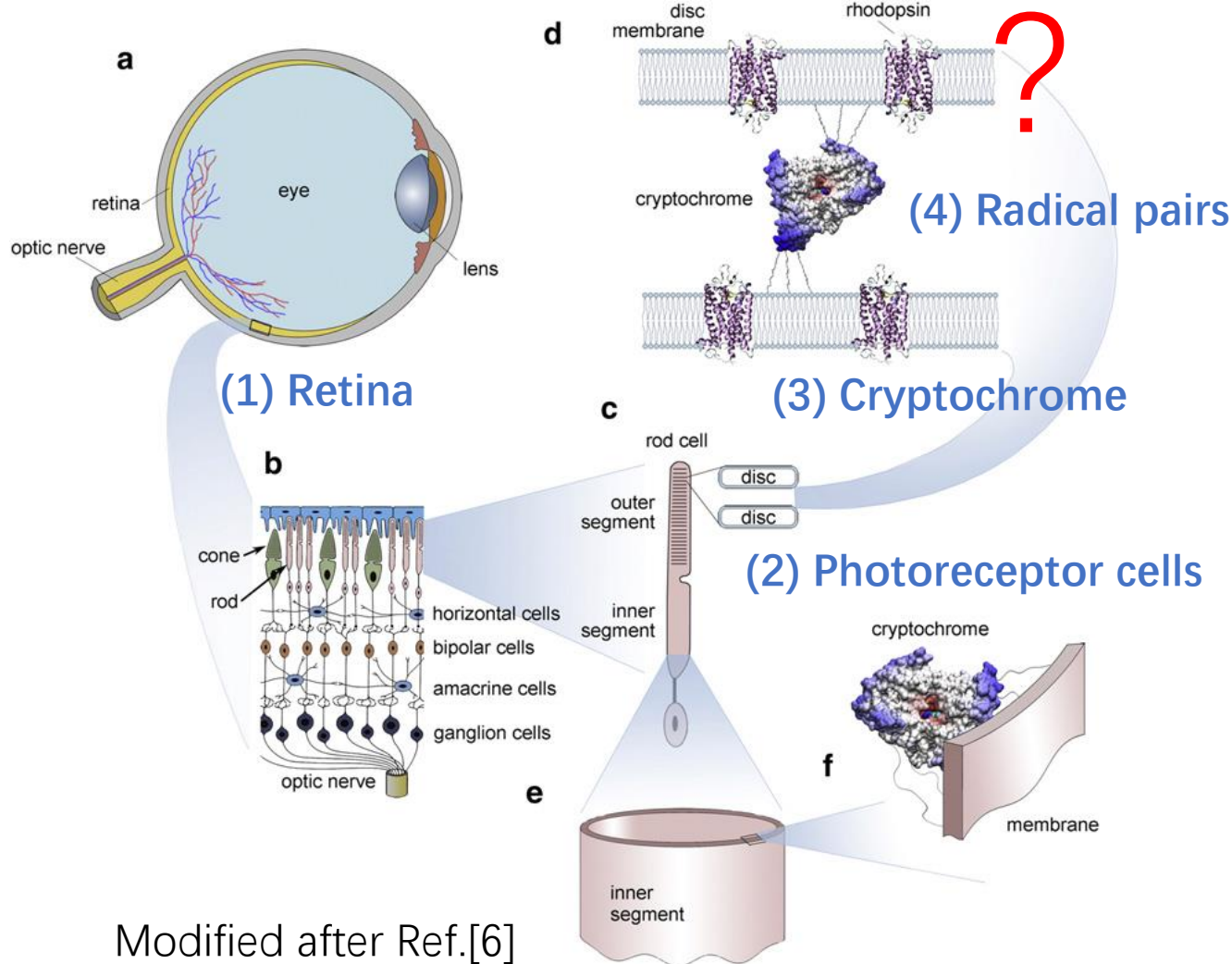
Modified after Ref.[3]



# Radical based magnetoreception

Magnetic information -> Chemical signal -> Electric signal

## (5) Visual pathway



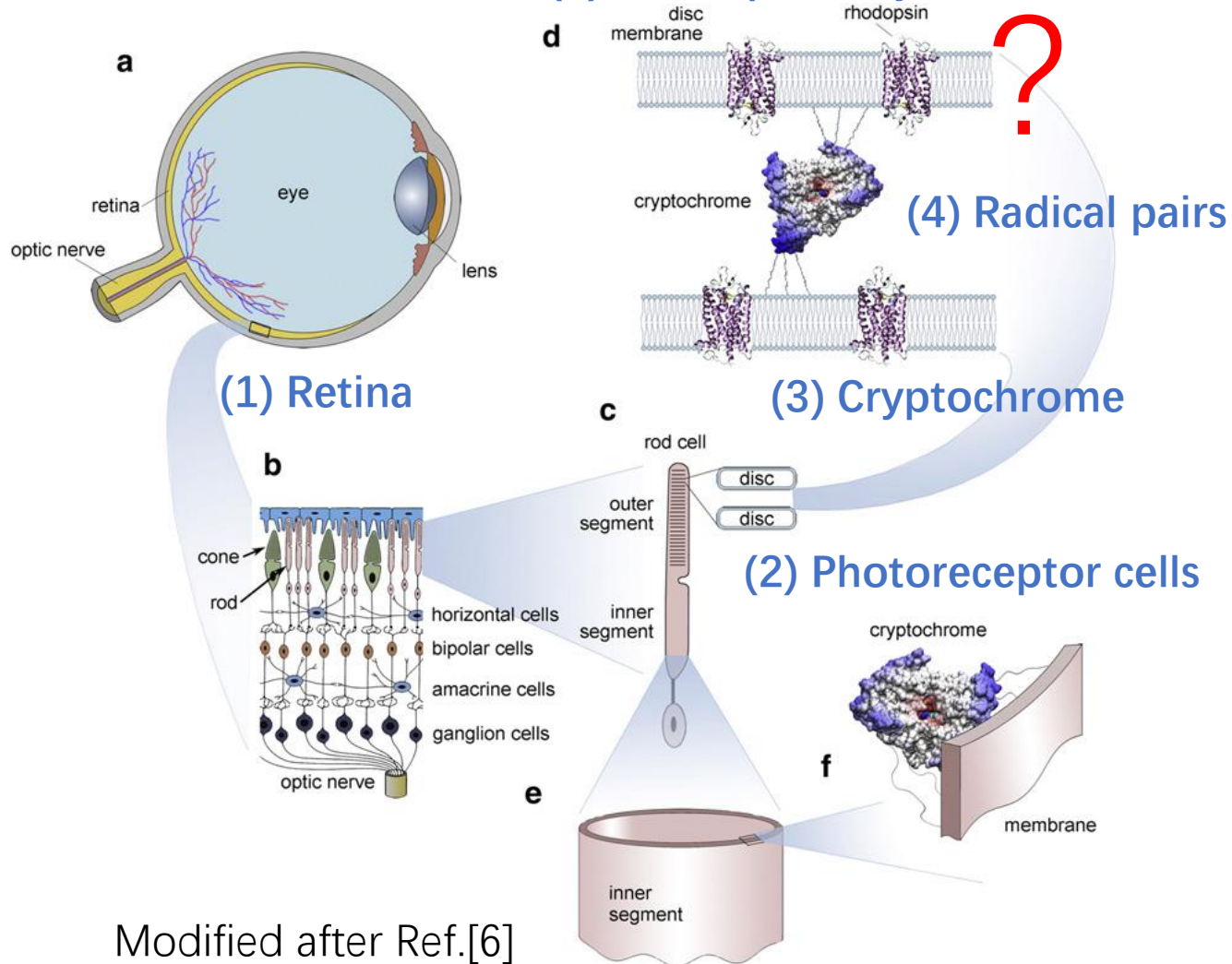
- 1) A **neural connection** between night vision and magnetic sensing in European robin.[12]
- 2) Different wavelength of incident light can **disrupt magnetic orientation** of European robin.[13,14]
- 3) The magnetic compass of European robin is **a inclination compass with <5° precision**, which can be explained by radical pair mechanism instead of any other hypothesis.[15,16]
- 4) Cryptochrome is the **only protein family** that can form **a light-induced radical pair**.

Modified after Ref.[6]

# Radical based magnetoreception

Magnetic information -> Chemical signal -> Electric signal

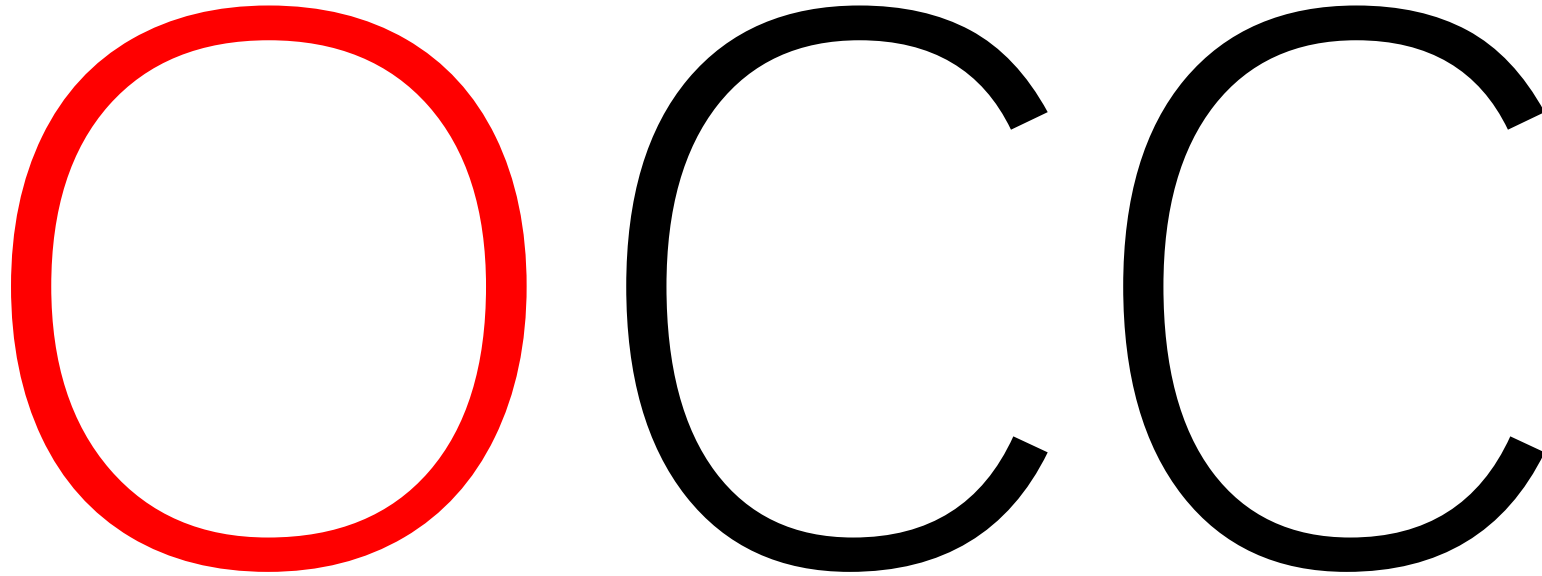
## (5) Visual pathway



Modified after Ref.[6]

- 5) **Cry4 protein**, a member of cryptochrome family, is found recently located on the **outer segment** in cones in birds' retina.[17]
- 6) A popular hypothesis of (4)-(5) is an noncanonical biological pathway of visual system that can modulate night vision. **No such signaling pathway** is found till now.
- 7) We propose that no downstream signaling involved and instead a photon competition plays a key role. A novel biophysical model:

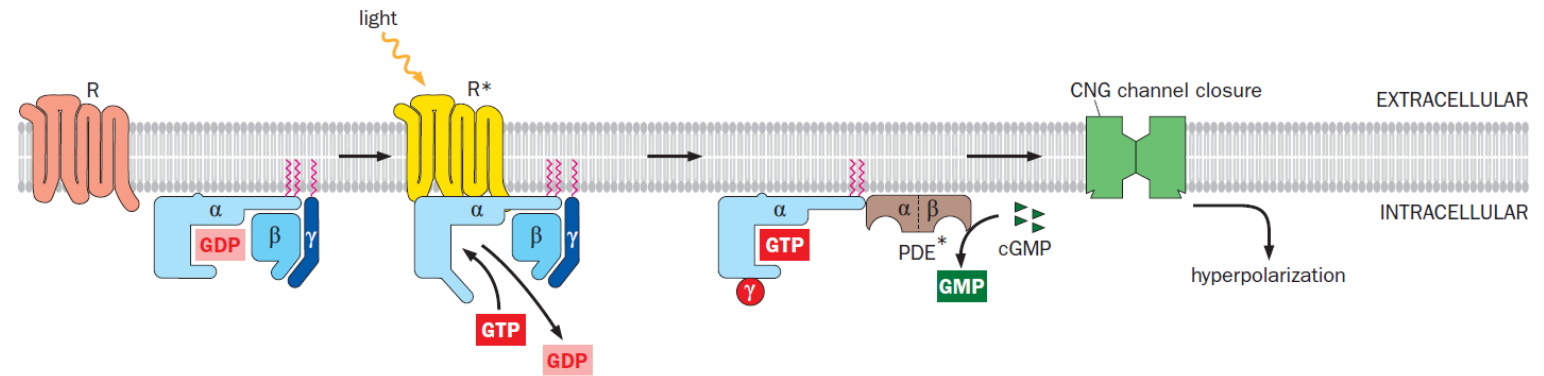
**Opsin-Cryptochrome competition model(OCC model).**



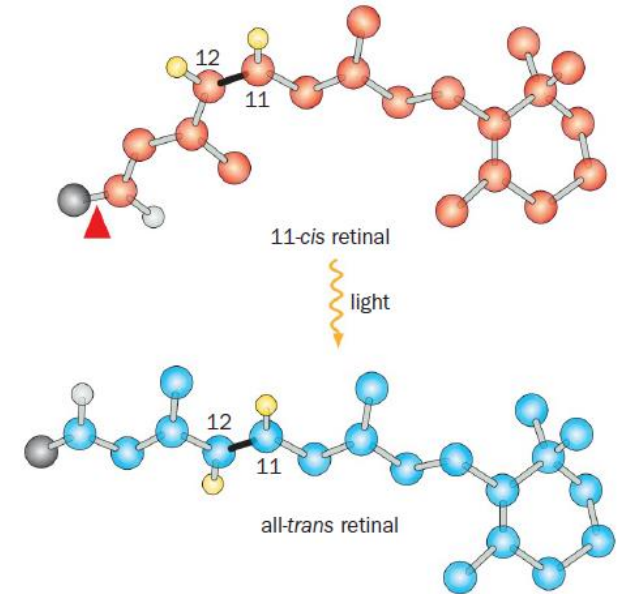
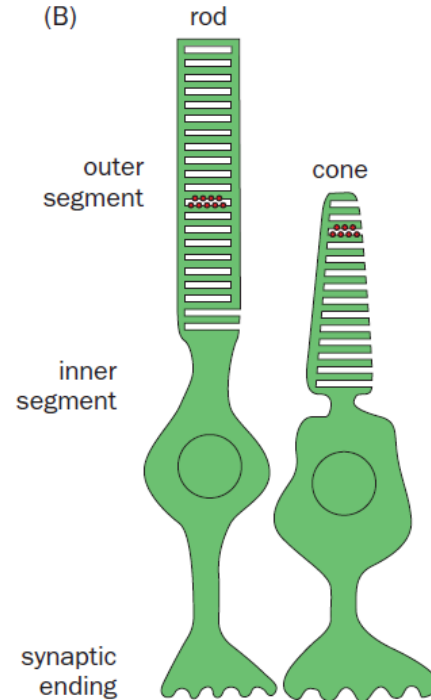
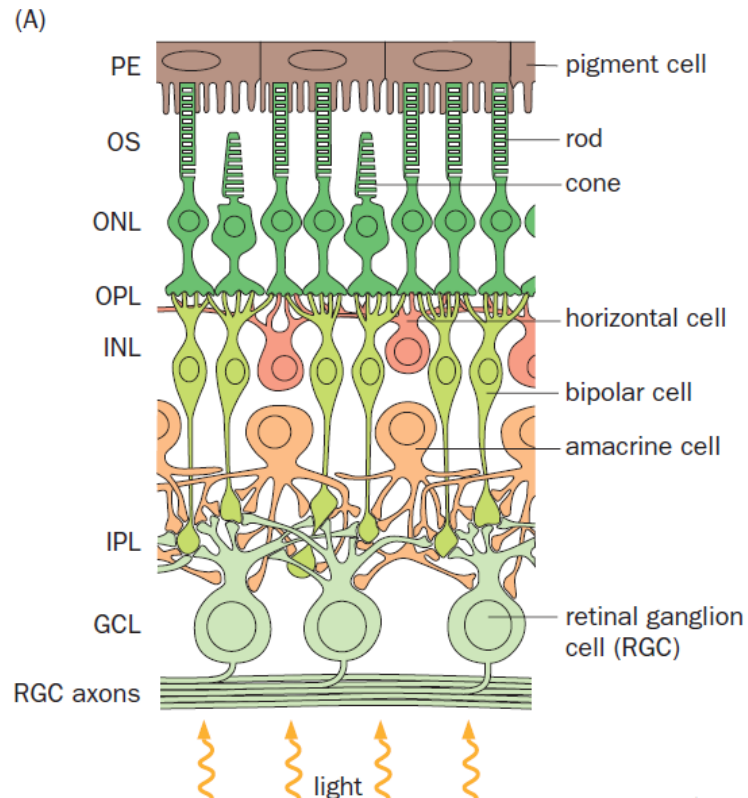
Opsin, Visual system



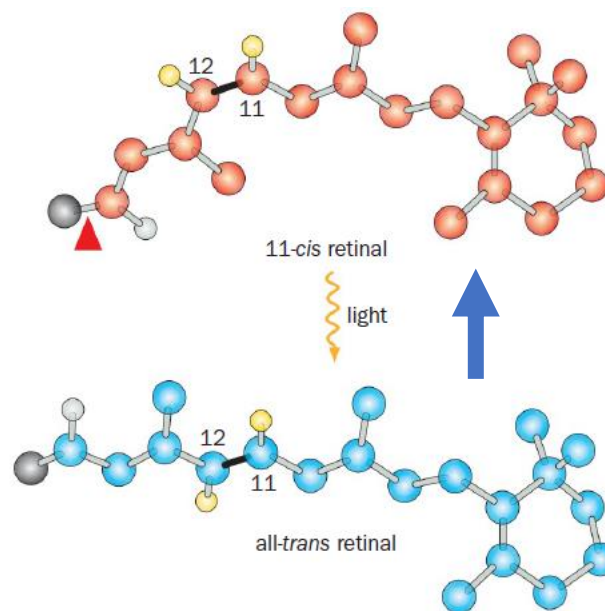
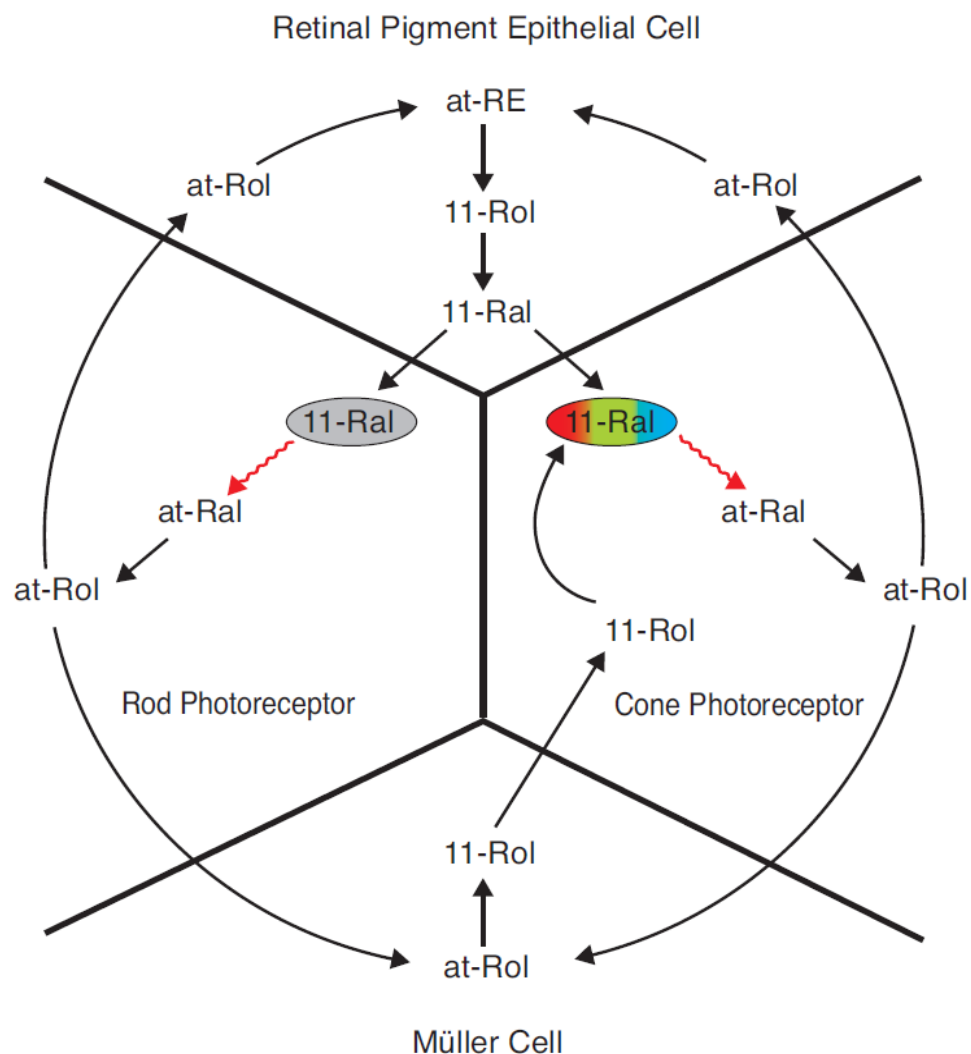
# Light perception



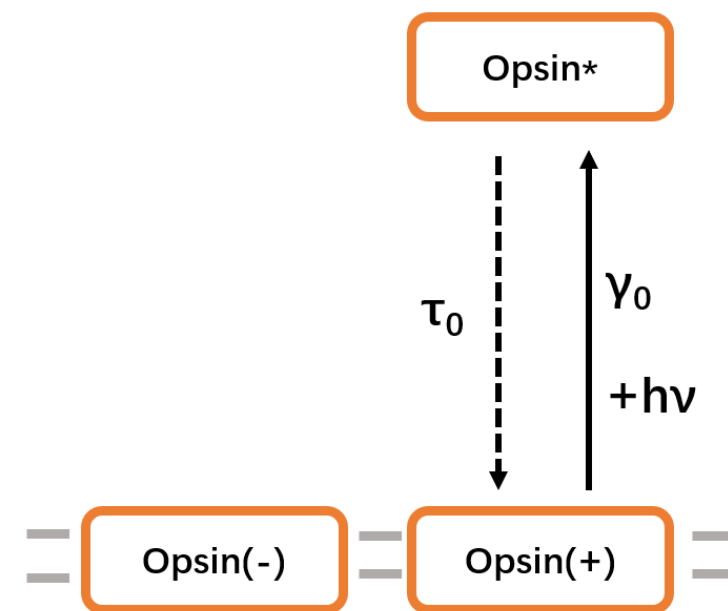
Phototransduction cascade



# A cone visual cycle: cis-retinal $\leftrightarrow$ trans-retinal

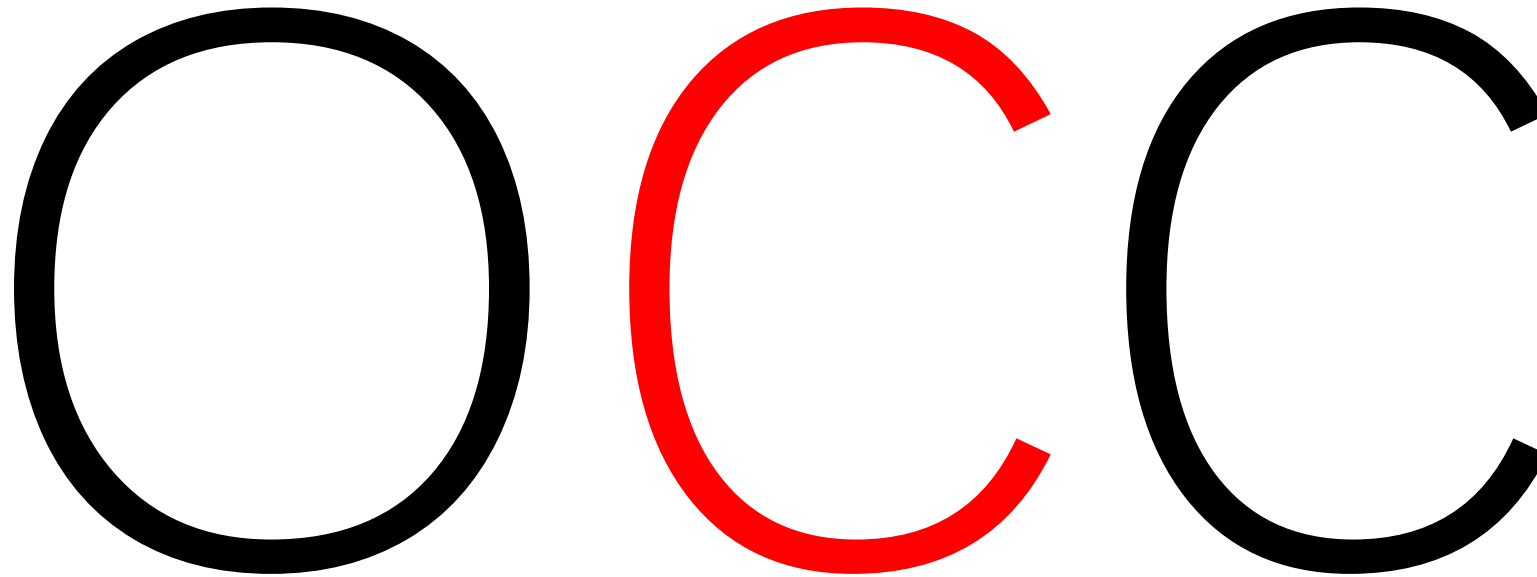


Retinal in a rhodopsin



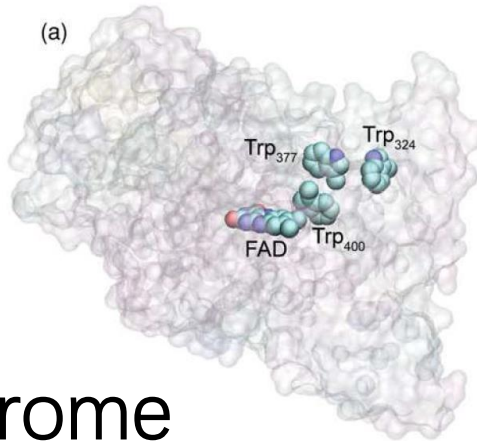
**Abbreviations:** at-RE, *all-trans*-retinyl esters; at-Ral, *all-trans*-retinal; at-Rol, *all-trans*-retinol; 11-Ral, 11-*cis*-retinal; 11-Rol, 11-*cis*-retinol.

Modified after Ref.[7,8]



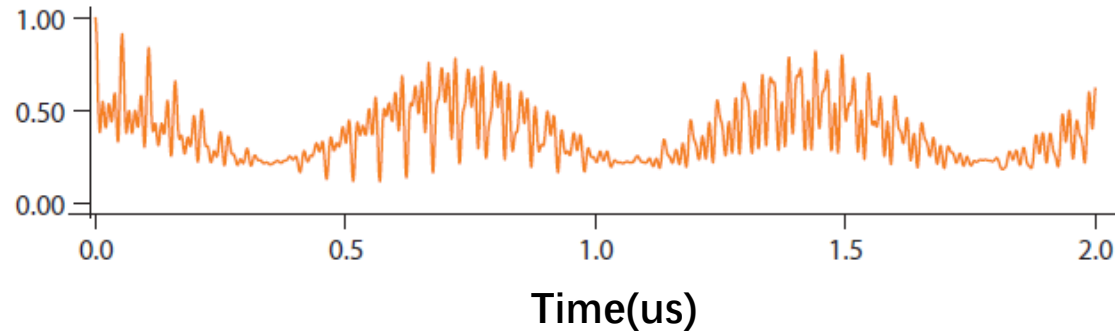
Cryptochrome, Radical pairs

# Principle of radical pair mechanism

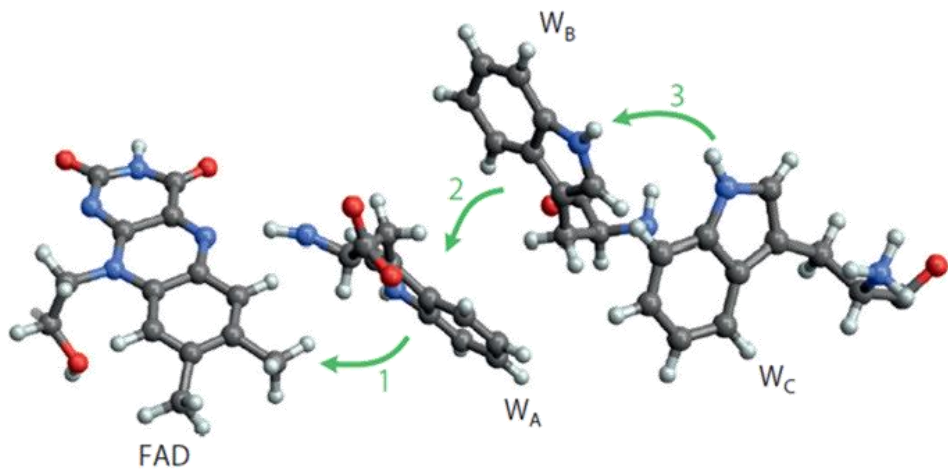


Cryptochrome

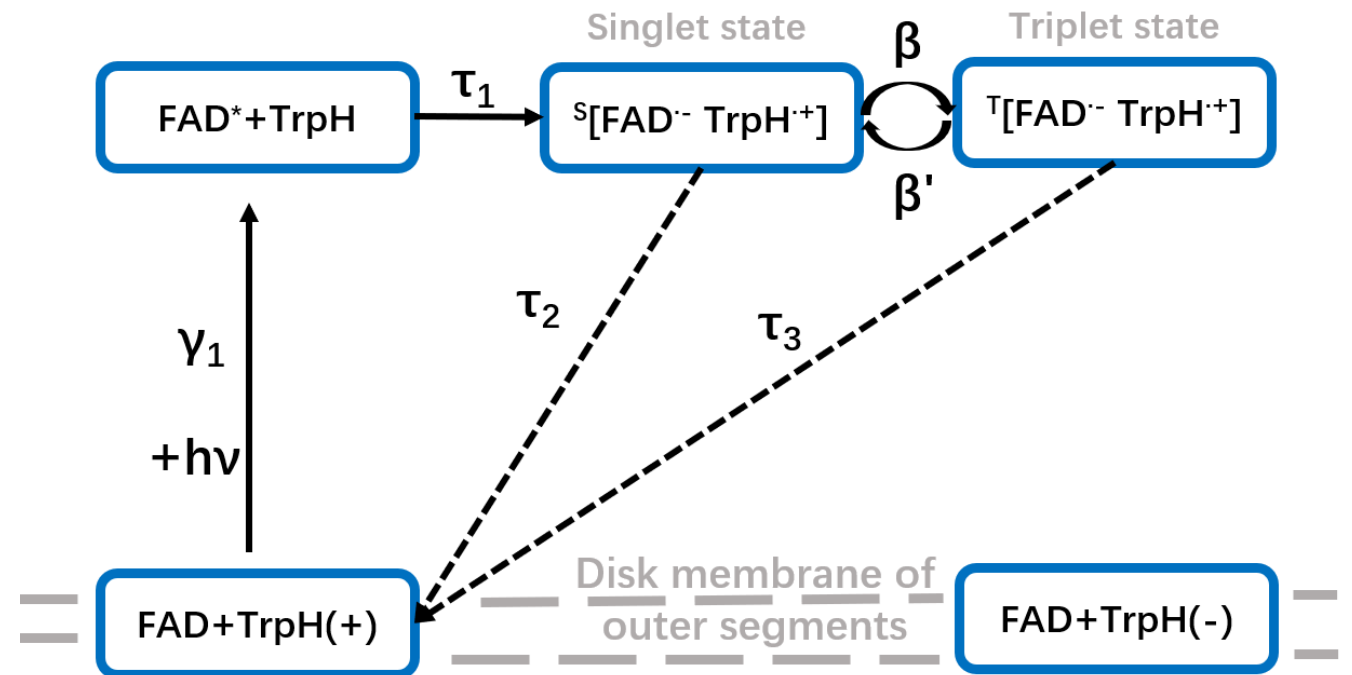
Singlet Fraction



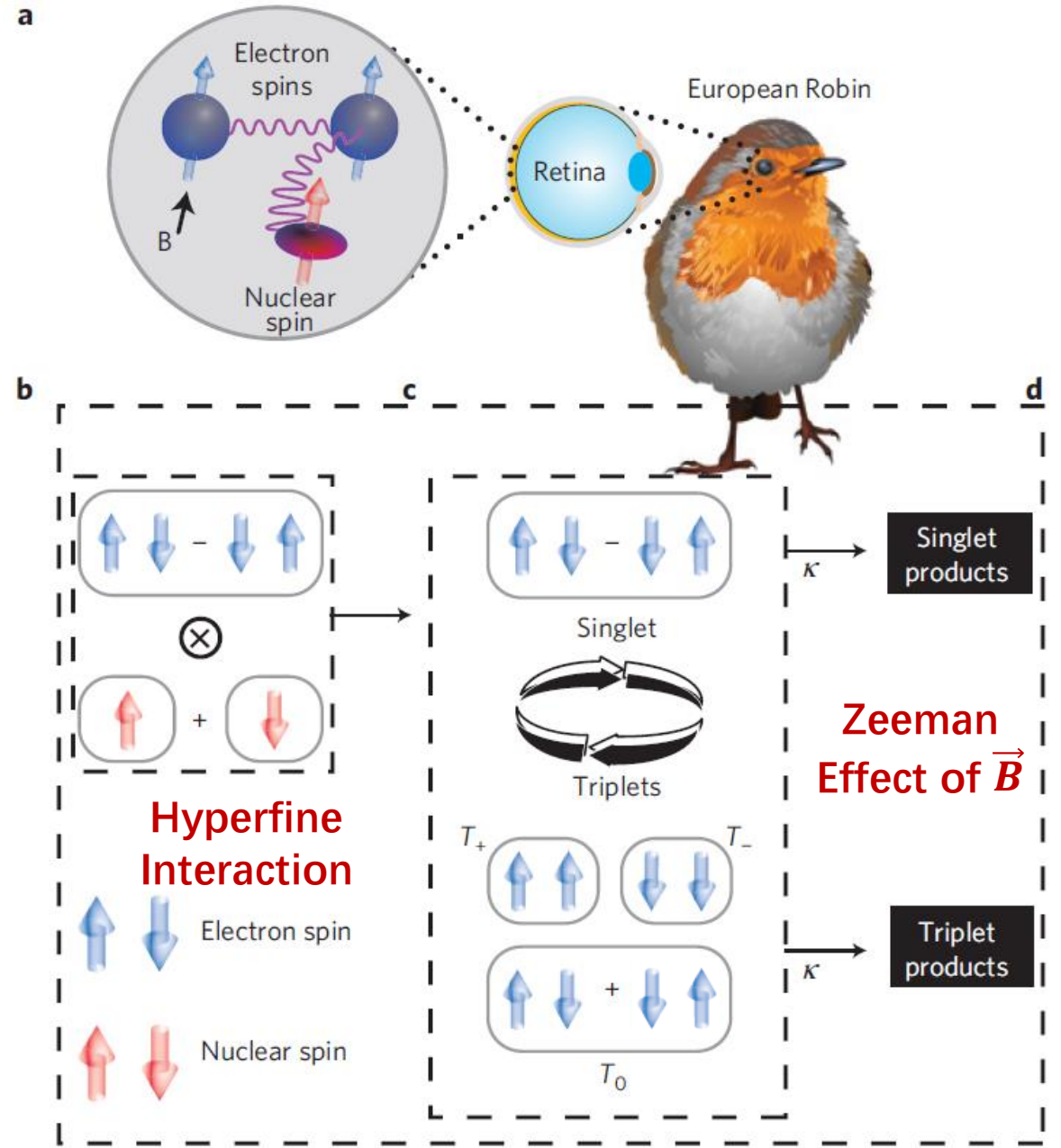
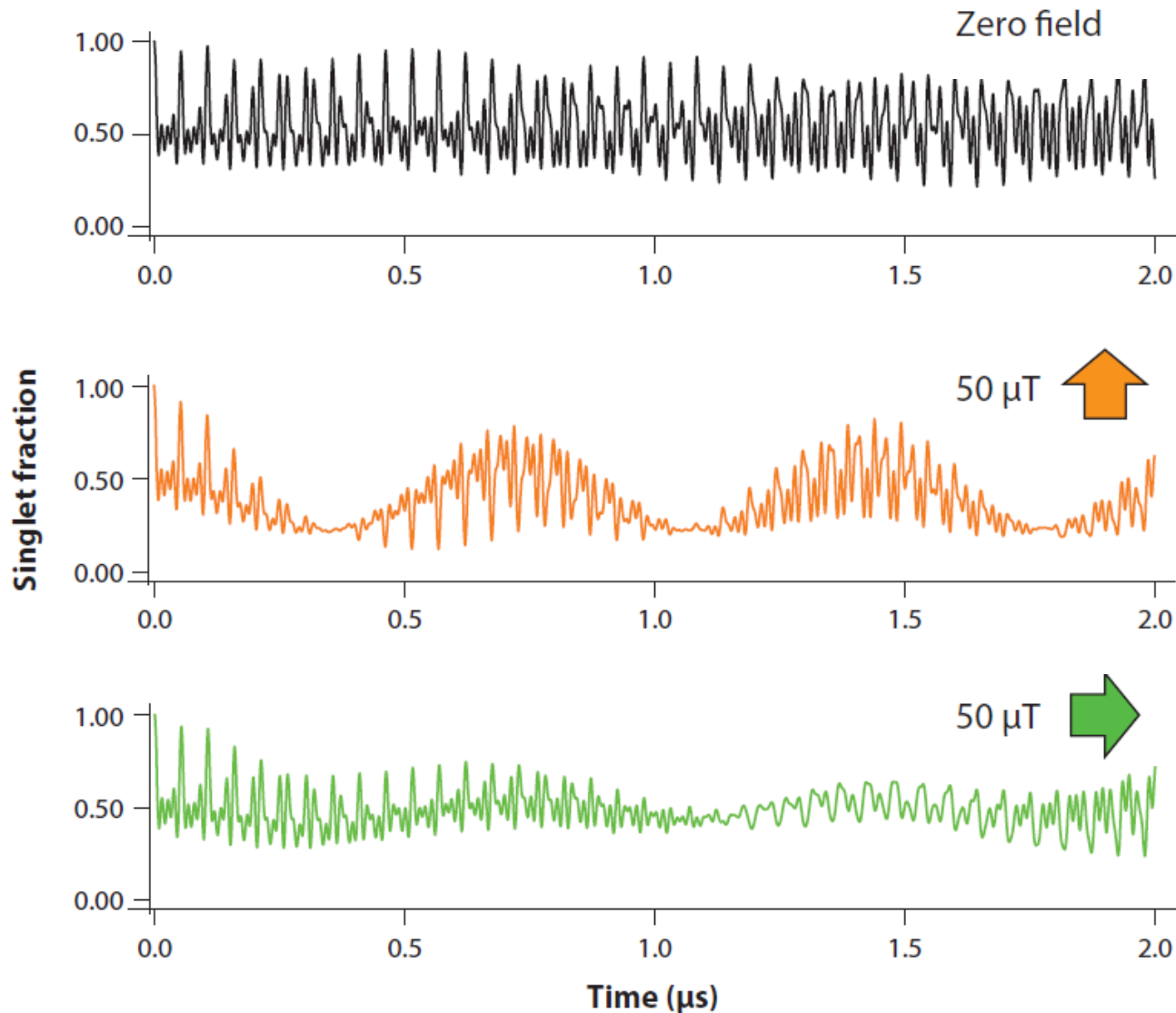
50  $\mu$ T



FAD - Flavin adenine dinucleotide  
Trp(W) - Tryptophan

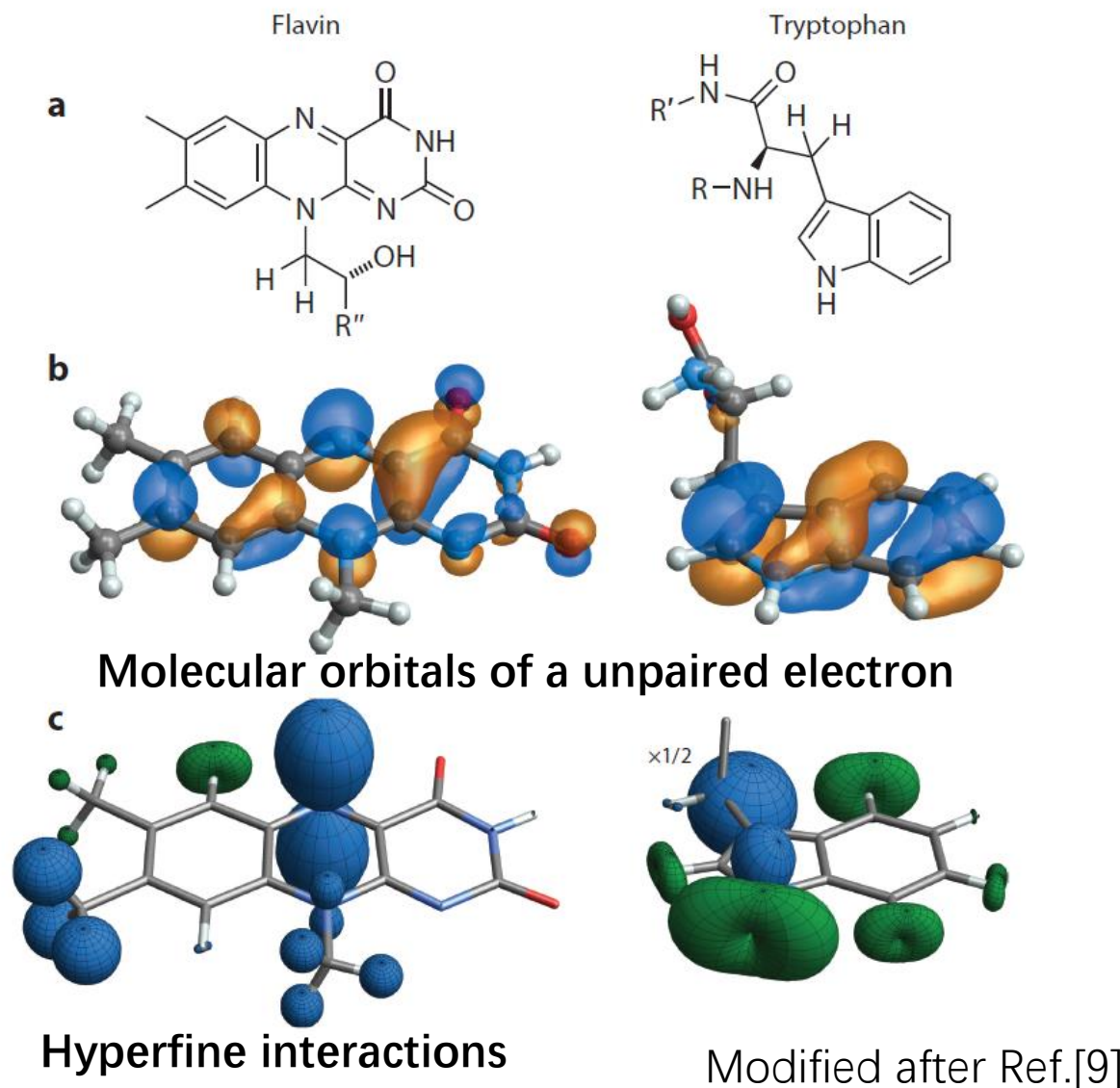


# Spin dynamics





# Quantum effect



The fractional yield of singlet and triplet state product is  $\phi_S = 1 - \phi_T$  is an integral of the real part of  $T(t)$  which is the singlet fraction at arbitrary time  $t$ .

$$\phi_T = k_T \int_0^\infty T(t) dt$$

$$T(t) = \text{Tr}[Q^T \rho(t)]$$

$Q^T$  is the projection operator of triplet state while  $\rho(t)$  is the density matrix of the radical pair at arbitrary time. The triplet state fraction is a trace operation of the density matrix projected on the triplet state product.

$$\rho(t) = \frac{1}{N} e^{-\frac{iHt}{\hbar}} \rho(0) e^{\frac{iHt}{\hbar}}$$

Assume there is only singlet product at time 0, where  $\rho(0) = Q^S e^{-kt}$ , here to make it simple, we let  $k = k_S = k_T$ .  $N$  is the number of nuclear spin.

The hamiltonian for two electron spins are  $H_1$  and  $H_2$  respectively.  $\vec{B}$  is the magnetic field vector with a specific direction.

$$H = H_1(\vec{B}) + H_2(\vec{B})$$

$$H_j = g\mu_B \vec{S}_j \cdot (\vec{B} + A_j \vec{I}_j)$$

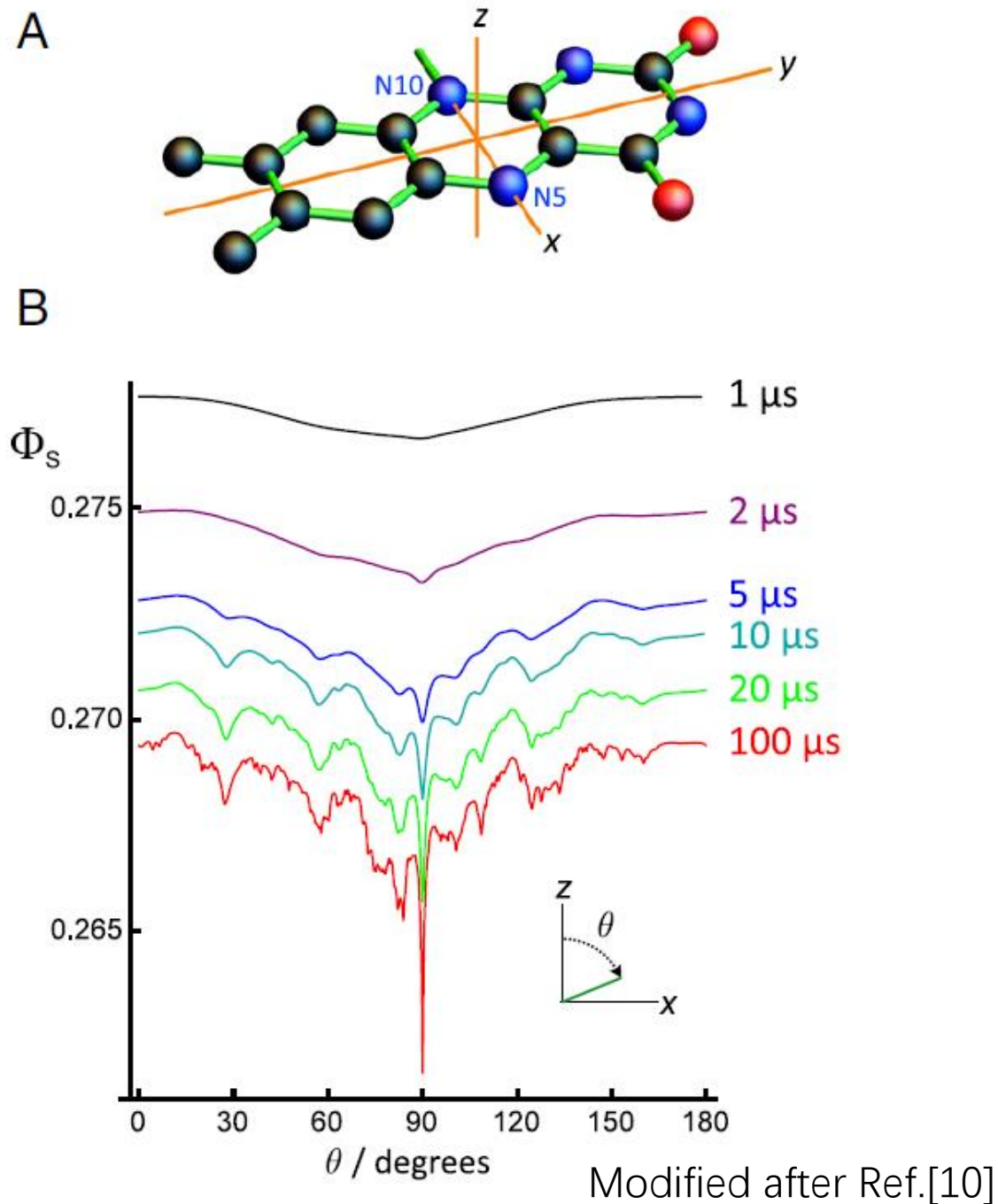
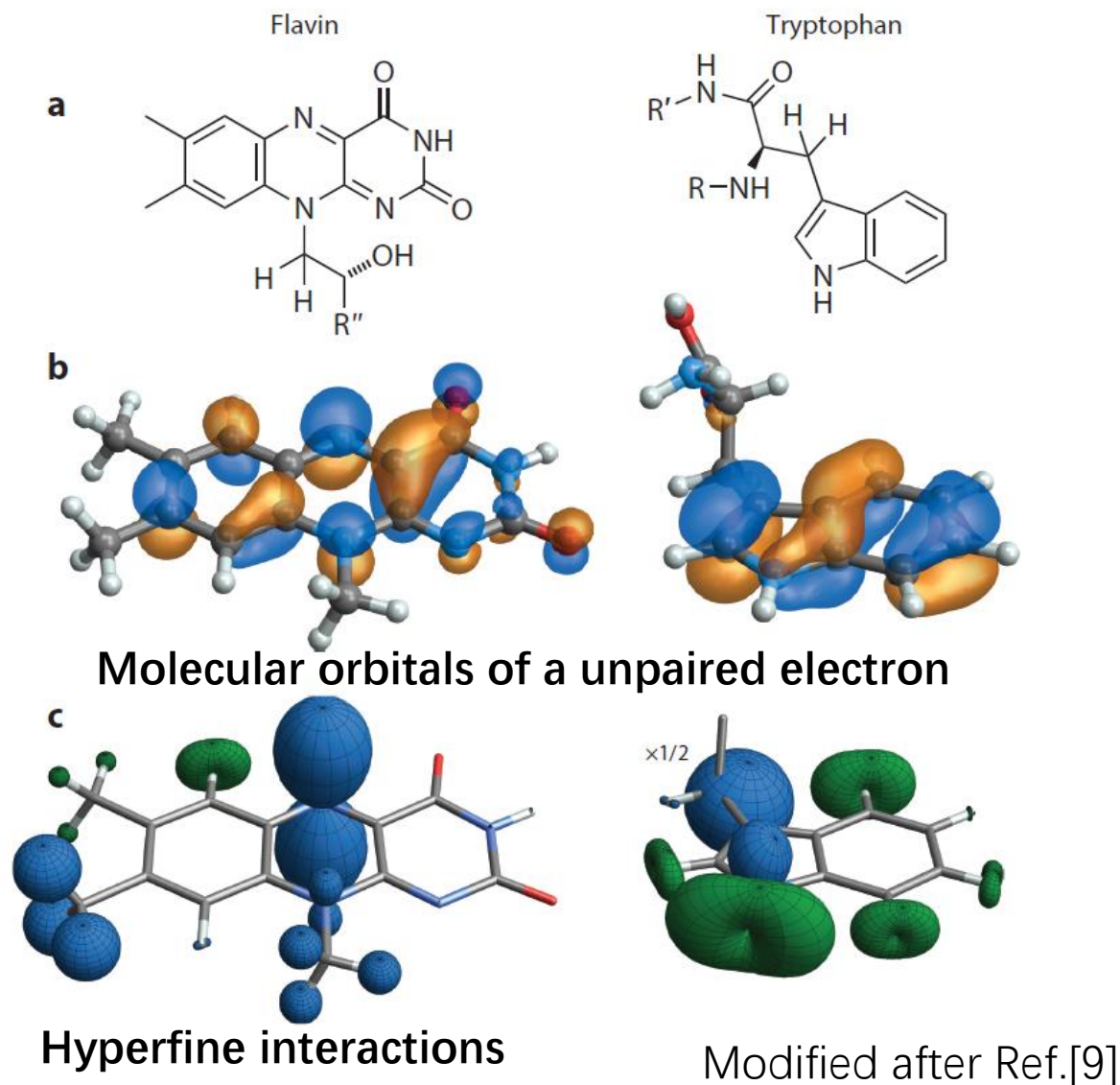
$\vec{S}_j$  is the electron spin operator whereas  $\vec{I}_j$  is the nuclear spin operator correlated with the nuclear spin in vicinity.  $A_j$  is the anisotropic coefficient of the nuclear spin.

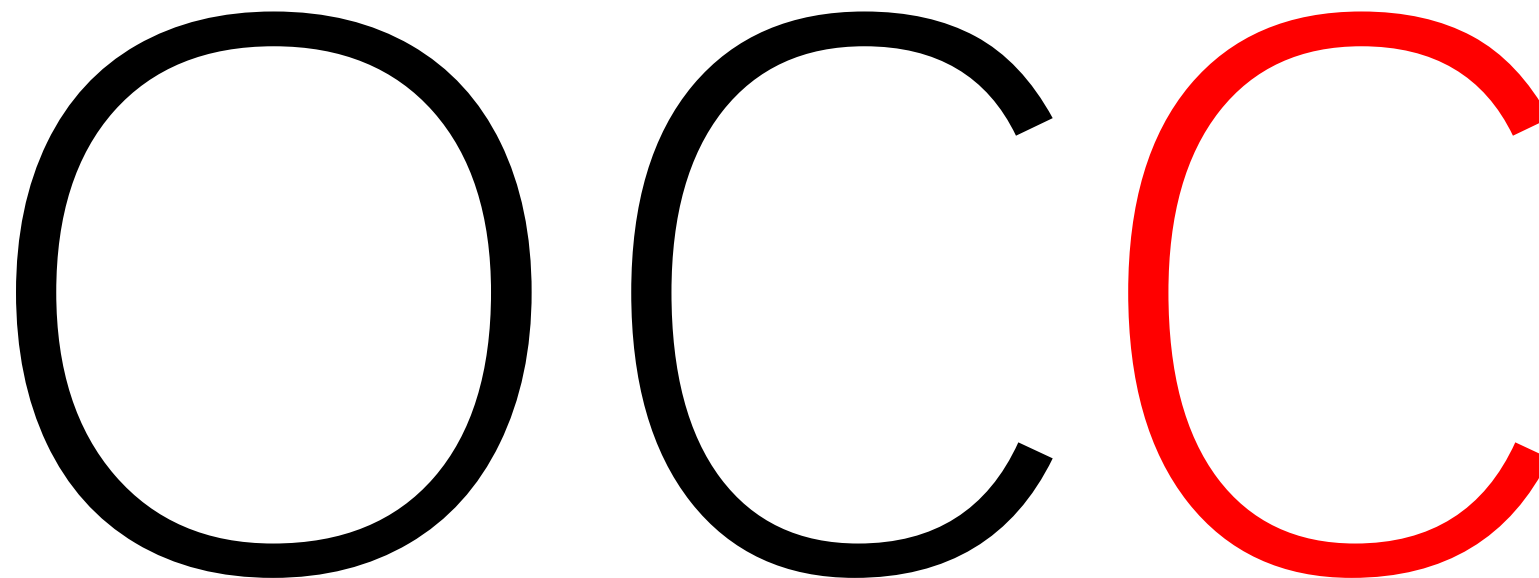
$$T(t) = \phi_S = 1 - \text{Tr}[Q^T \frac{1}{N} e^{-\frac{iHt}{\hbar}} Q^S e^{-kt} e^{\frac{iHt}{\hbar}}]$$

$$\phi_S = 1 - \frac{1}{N} \sum_{mn} Q_{mn}^T Q_{nm}^S \frac{k^2}{k^2 + (\omega_m - \omega_n)^2}$$



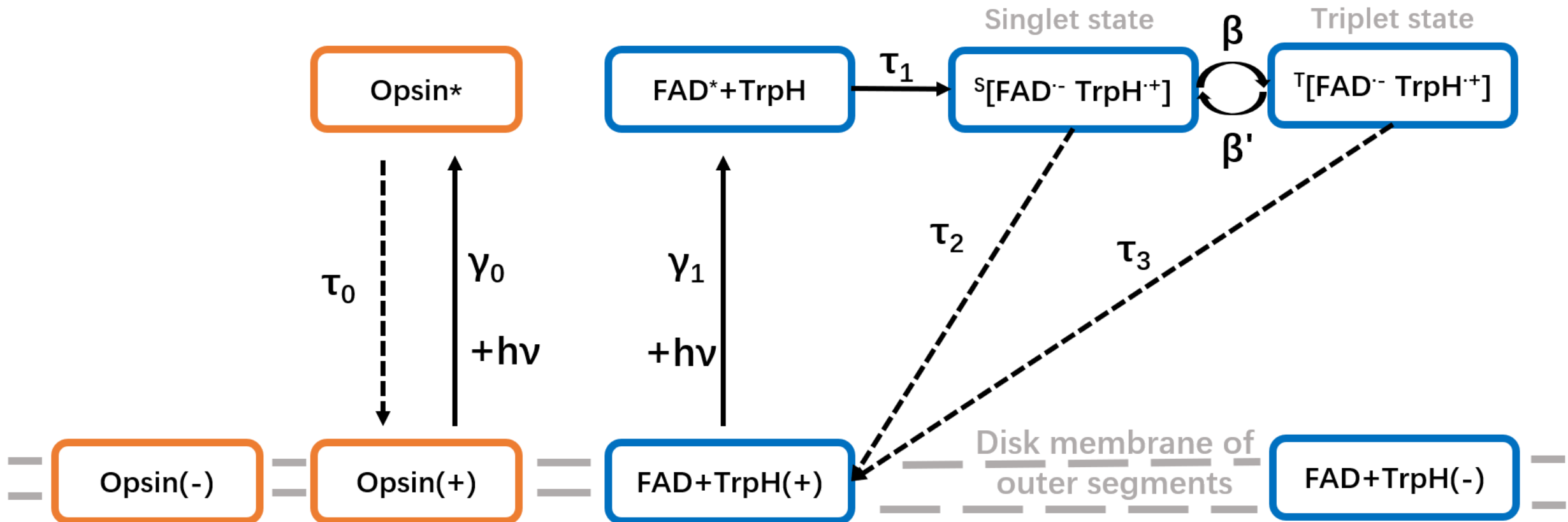
# Quantum effect





Photon Competition

# Opsin-Cryptochrome Competition model

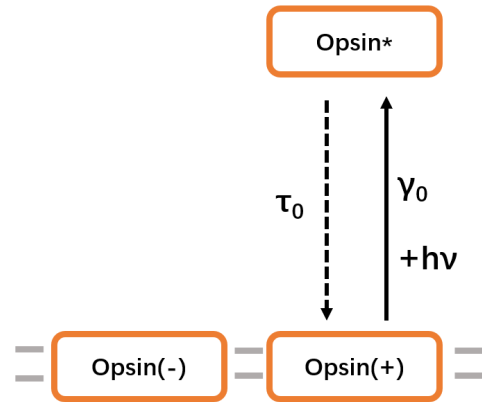


(+) represents the ground state of a molecule absorbed by one photon; (\*) represents the excitation state of a molecule after photon absorption; (s) and (t) represent the singlet and triplet state of a molecule.

# Model formulation

$$N_{op}^{(+)} + N_{FAD}^{(+)} = N_{photon} \quad (9)$$

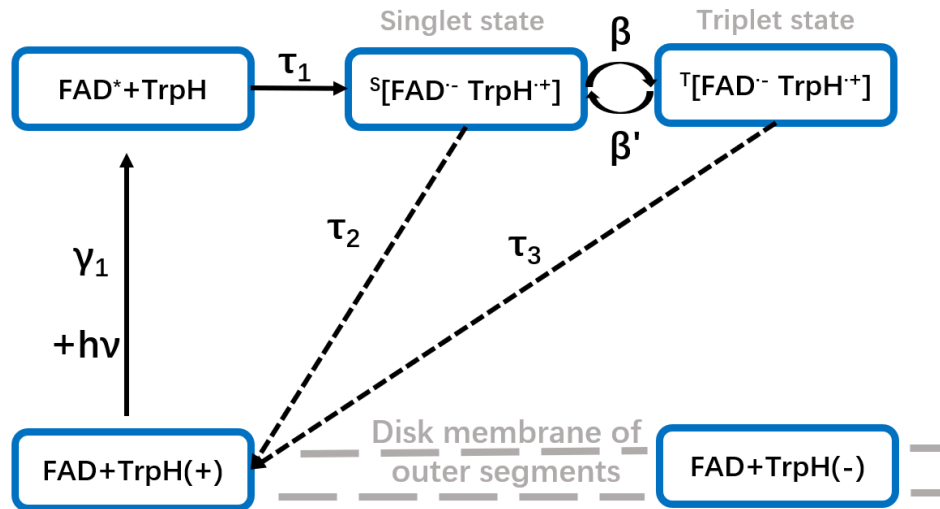
$$R_{opsin} = \frac{N_{op}^{(+)}}{N_{photon}} \quad (10)$$



$$N_{op}^{(+)} + N_{op}^* + N_{op}^{(-)} = C_1 \quad (1)$$

$$\frac{d(N_{op}^{(+)} + N_{op}^{(-)})}{dt} = -\gamma_0 N_{op}^{(+)} + \frac{1}{\tau_0} N_{op}^* \quad (2)$$

$$\frac{dN_{op}^*}{dt} = \gamma_0 N_{op}^{(+)} - \frac{1}{\tau_0} N_{op}^* \quad (3)$$



$$N_{FAD}^{(+)} + N_{FAD}^* + N_{FAD}^{Singlet} + N_{FAD}^{Triplet} + N_{FAD}^{(-)} = C_2 \quad (4)$$

$$\frac{d(N_{FAD}^{(+)} + N_{FAD}^{(-)})}{dt} = -\gamma_1 N_{FAD}^{(+)} + \frac{1}{\tau_2} N_{FAD}^{Singlet} + \frac{1}{\tau_3} N_{FAD}^{Triplet} \quad (5)$$

$$\frac{dN_{FAD}^*}{dt} = \gamma_1 N_{FAD}^{(+)} - \frac{1}{\tau_1} N_{FAD}^* \quad (6)$$

$$\frac{dN_{FAD}^{Singlet}}{dt} = \frac{1}{\tau_1} N_{FAD}^* - \frac{1}{\tau_2} N_{FAD}^{Singlet} - \beta N_{FAD}^{Singlet} + \beta' N_{FAD}^{Triplet} \quad (7)$$

$$\frac{dN_{FAD}^{Triplet}}{dt} = \beta N_{FAD}^{Singlet} - \beta' N_{FAD}^{Triplet} - \frac{1}{\tau_3} N_{FAD}^{Triplet} \quad (8)$$

A stationary solution: let all derivatives=0

$$N_{op}^{(+)} + N_{FAD}^{(+)} = N_{photon} \quad (9)$$

$$R_{opsin} = \frac{N_{op}^{(+)}}{N_{photon}} \quad (10)$$

$$N_{op}^{(-)} + B * N_{op}^{(+)} = C_1 \quad (11)$$

$$N_{FAD}^{(-)} + A * N_{FAD}^{(+)} = C_2 \quad (12)$$

$$(A - B)N_{op}^{(+)^2} + (C_1 + C_2 - N_{photon}(A - B))N_{op}^{(+)} - N_{photon}C_1 = 0 \quad (15)$$

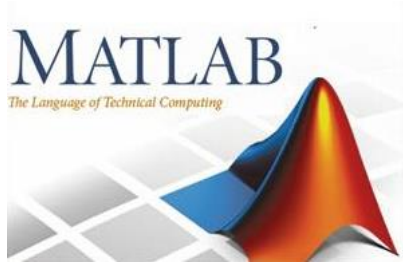
$$R_{opsin} = \frac{-(C_1 + C_2 - N_{photon}(A - B)) + \sqrt{(C_1 + C_2 - N_{photon}(A - B))^2 + 4N_{photon}C_1(A - B)}}{2N_{photon}(A - B)} \quad (16)$$

$$A = 1 + \gamma_1 \left( \tau_1 + \frac{1}{\frac{\beta'}{\tau_2\beta} + \frac{1}{\tau_2\tau_3\beta} + \frac{1}{\tau_3}} + \frac{1}{\frac{1}{\tau_2} + \frac{\beta}{\tau_3\beta' + 1}} \right) \quad B = \tau_0\gamma_0 + 1$$

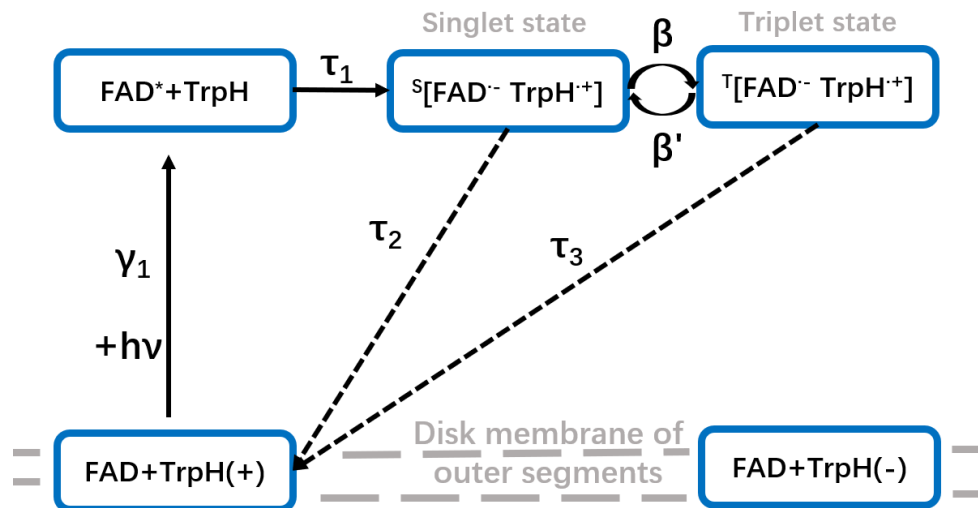
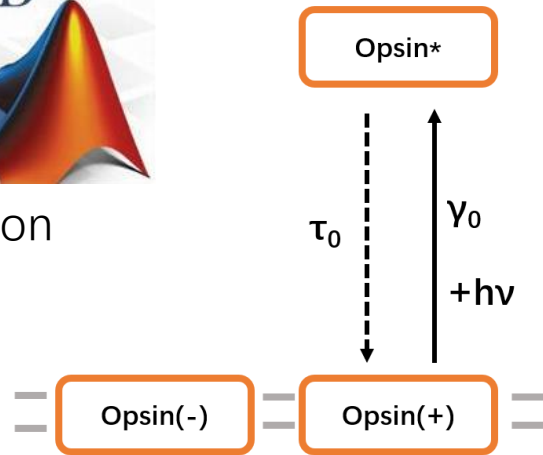
# Simulation



# Parameter initiation



Simulation



- **Singlet-triplet oscillation  $\beta$  coefficient:**

$$\beta' = 0; \beta = \frac{\Delta(\Phi T)}{\Delta(\tau_2)} \text{ in terms of angle}$$

- **Reciprocal of rate constant(lifetime):**

$$\tau_0 = 4.2\text{ms}; \tau_1 = 1\text{ns};$$

$$\tau_2 = 1\mu\text{s}; \tau_3 = 10\text{ms};$$

- **Rate constant of light absorption:**

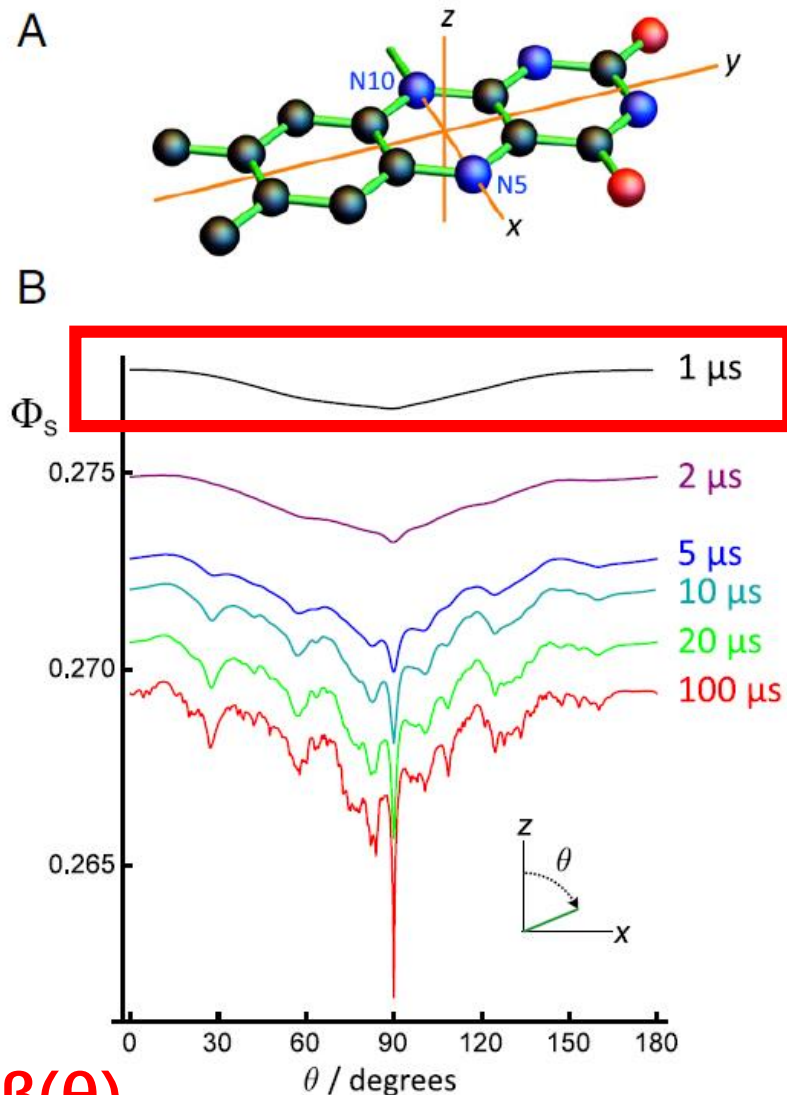
$$\gamma_0 = \gamma_1 = 10^9/\text{s};$$

- Assume the **protein number** of opsin and cryptochrome is the same:  $C_1 = C_2$

- **Incident photon number**  $N_{\text{photon}}$

$$\text{where } N_{\text{photon}} : C_1 = 10^{-3}$$

# Parameter initiation

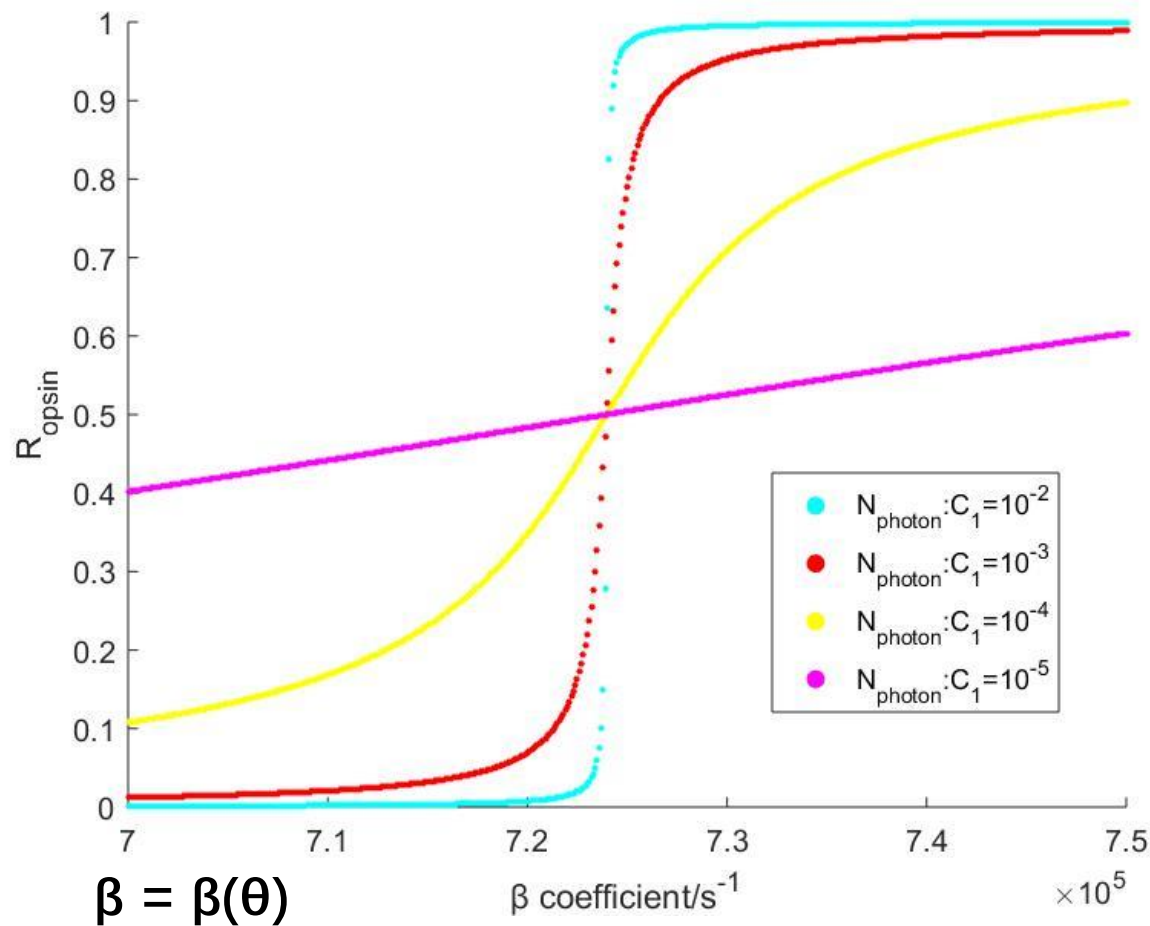


$$\beta = \beta(\theta)$$

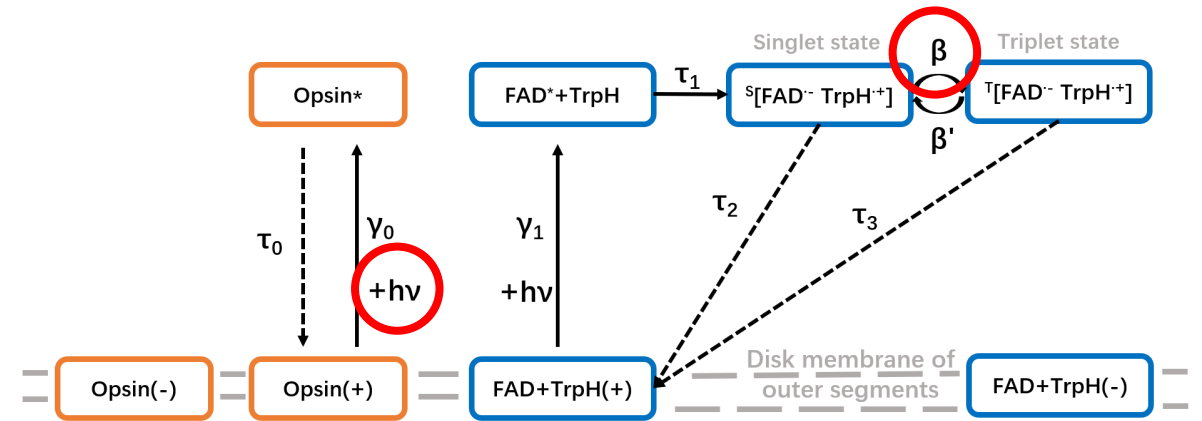
- Singlet-triplet oscillation  $\beta$  coefficient:  
 $\beta' = 0$ ;  $\beta = \frac{\Delta(\Phi T)}{\Delta(\tau_2)}$  in terms of angle
- Reciprocal of rate constant(lifetime):  
 $\tau_0 = 4.2\text{ms}$ ;  $\tau_1 = 1\text{ns}$ ;  
 $\tau_2 = 1\mu\text{s}$ ;  $\tau_3 = 10\text{ms}$ ;
- Rate constant of light absorption:  
 $\gamma_0 = \gamma_1 = 10^9/\text{s}$ ;
- Assume the protein number of opsin and cryptochrome is the same:  $C_1 = C_2$
- Incident photon number  $N_{\text{photon}}$   
 where  $N_{\text{photon}}:C_1 = 10^{-3}$

# Secondary amplification effect

$$R_{opsin} = \frac{-(C_1 + C_2 - N_{photon}(A - B)) + \sqrt{(C_1 + C_2 - N_{photon}(A - B))^2 + 4N_{photon}C_1(A - B)}}{2N_{photon}(A - B)}$$

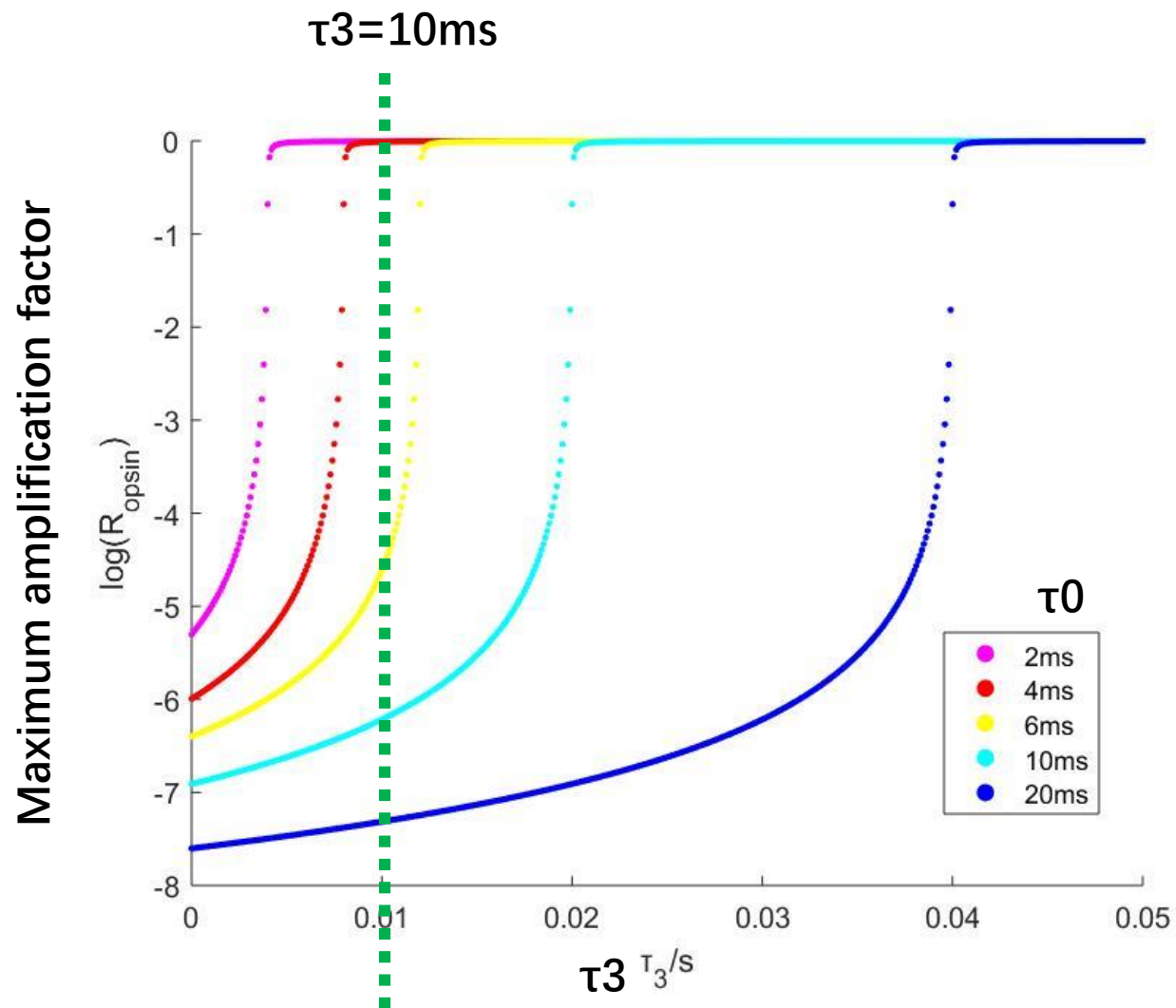


$R_{opsin} \in (0, 1)$  representing the percentage of incident photons that are absorbed by rhodopsin.

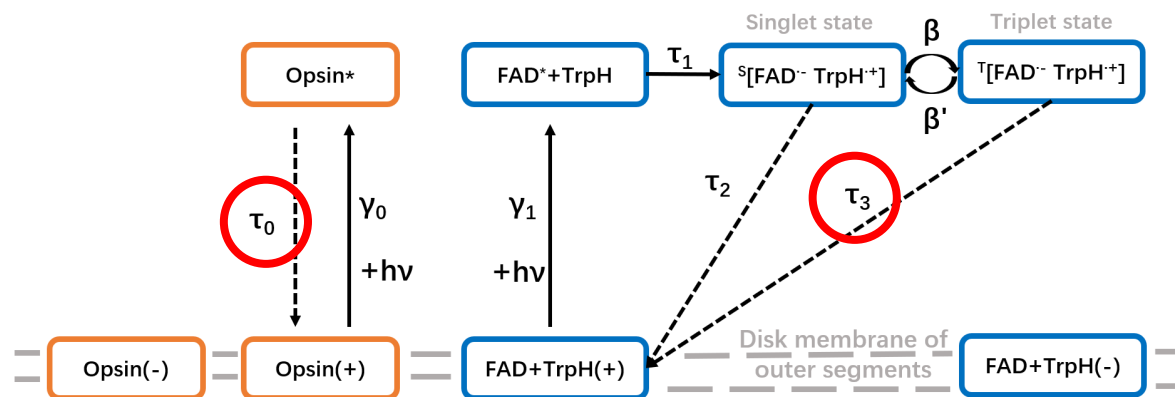


- Ratio of incident photons and protein numbers are critical for amplification **indicating a photon threshold** for magnetoreception.

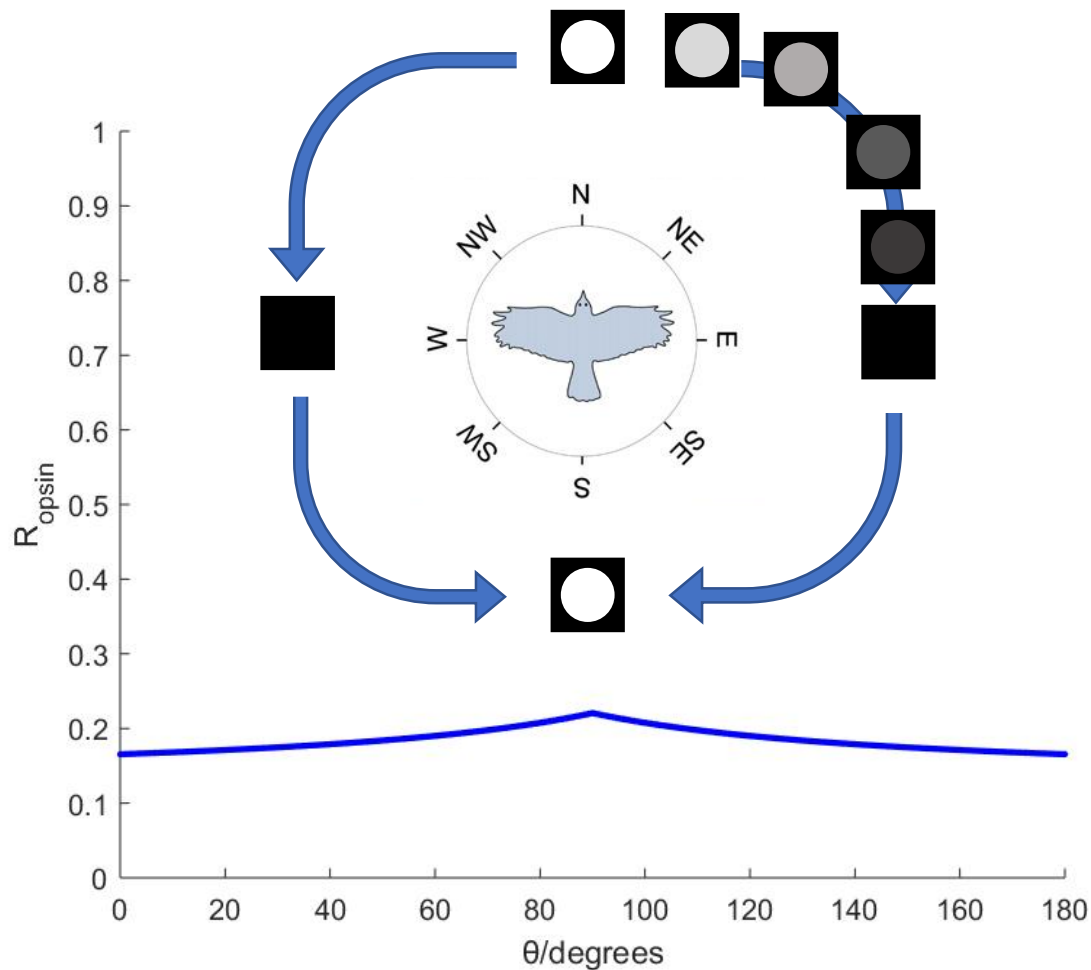
# Secondary amplification effect



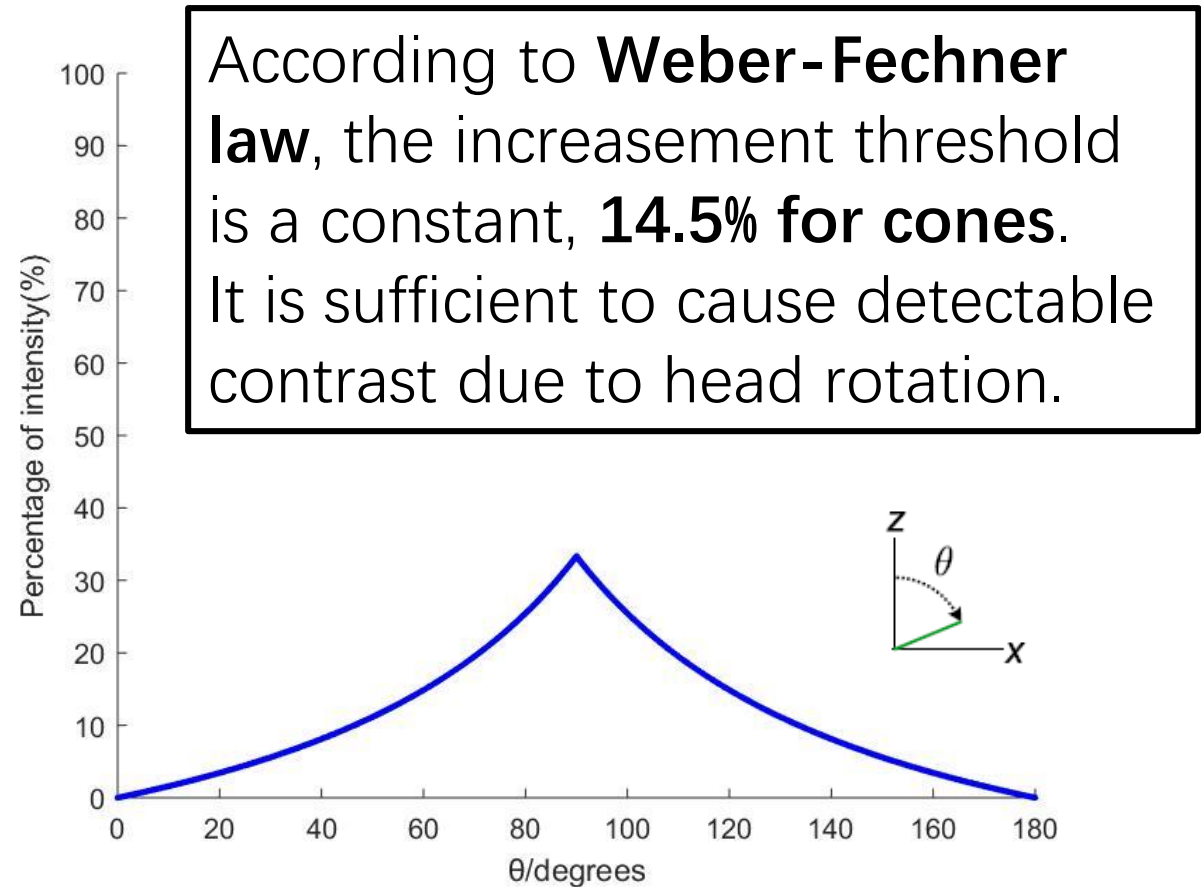
- The quantity of  $\tau_0$  is critical for the amplification effect.
- The secondary amplification effect is a key step to **convert a physical cause into biological consequence**.



# Photon competition simulation result



$$R_{opsin} = R_{opsin}(\beta) \text{ and } \beta = \beta(\theta)$$



$$\text{Incresement threshold} = \frac{\Delta I}{I}$$

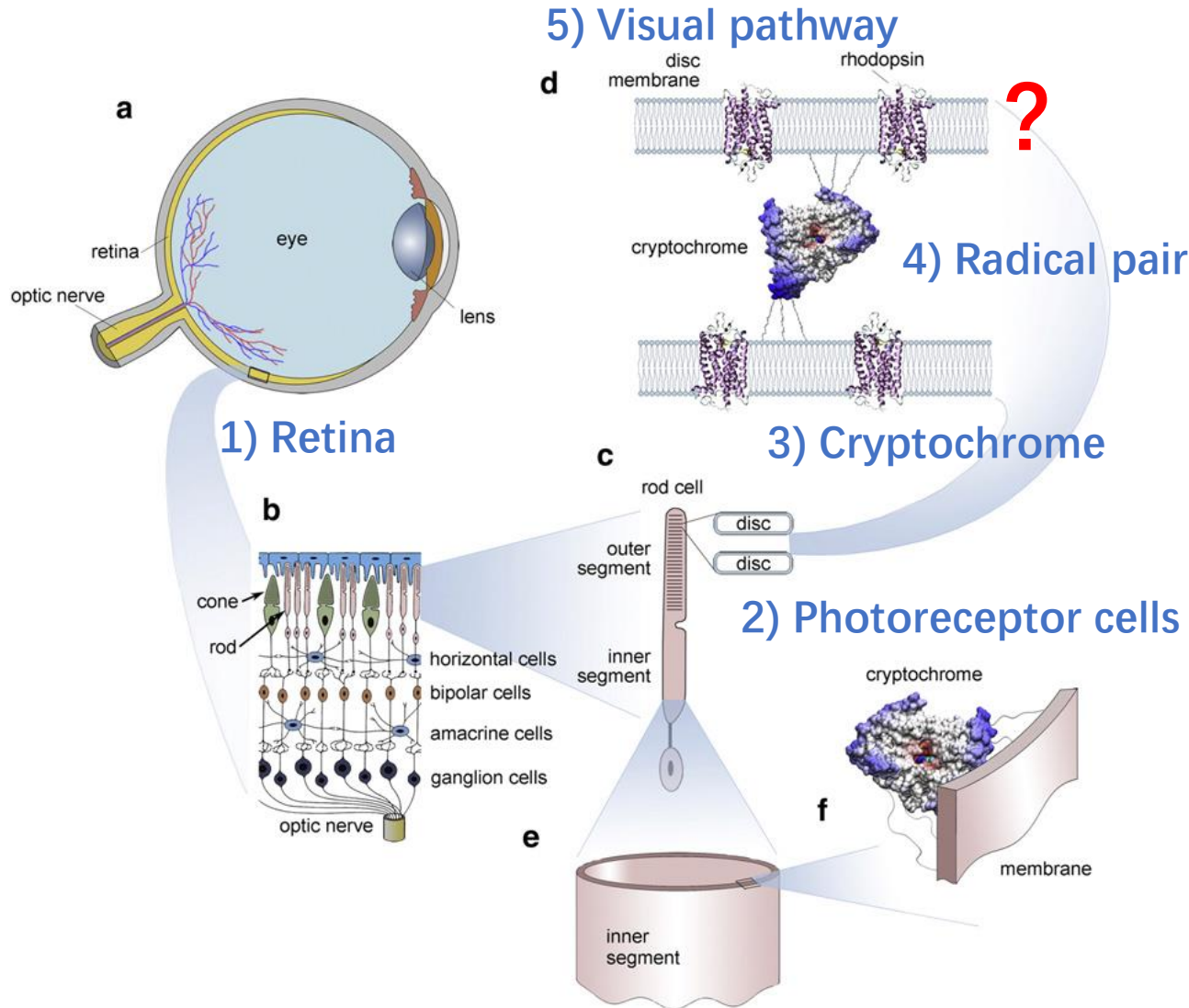
# Experimental predictions

- Two strategies to verify OCC model:
- (1) **Incident photon threshold** for magnetic sensing.
- There are lower and upper photon thresholds for competition capability of opsin and cryptochrome. Controlled light intensity of background can be a on-off effect of magnetoreception.
- (2) **Disruption/Analogy** of the magnetic correlated light pattern.
- Under zero magnetic field, generate an artificial light pattern resembling the earth magnetic field effect.
- Use changing spatial intensity of incident light on the retina to disrupt the existing light pattern caused by earth magnetic field .



# Summary

? ➡ OCC?



- A popular hypothesis of (4)-(5) is an noncanonical biological pathway of visual system that can modulate night vision. **No such signaling pathway** is found till now.
  - We propose that no downstream signaling involved and instead a photon competition plays a key role.
- A novel biophysical model:

**Opsin-Cryptochrome competition model(OCC model).**

Experiments to verify this model are proposed.

# Discussion

- Our model can not give a prediction whether **cones or rods or both are magnetic sensors**. One reason is that we don't have more precise parameters from experiments. The other is that **it is not fully understood** how rods and cones sense various light intensity.

Two key questions which we still know nothing about:

- **What is the evolutionary relationship among animals?**
- European robin, pigeon and turtle seems use a inclination magnetic compass whereas other animals use intensity or polarity compass.
- **How magnetic field is sensed and how the magnetic information is processed in different animal species?**
- Three possible mechanisms may account for magnetoreception of different animal species. But all of them are incomplete theories with a huge gap between a physical cause and a biological consequence.

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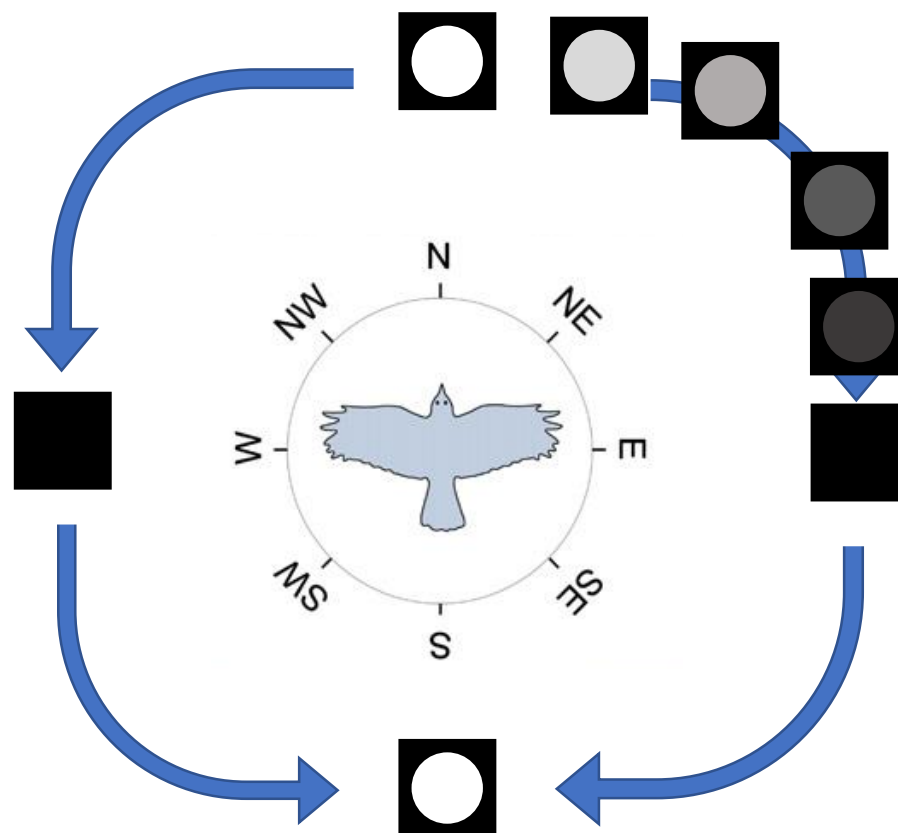
Jiansheng Wu,  
Assistance professor  
Department of Physics  
SUSTech



<https://github.com/LokyWei/Biophysical-model>



# Thank you

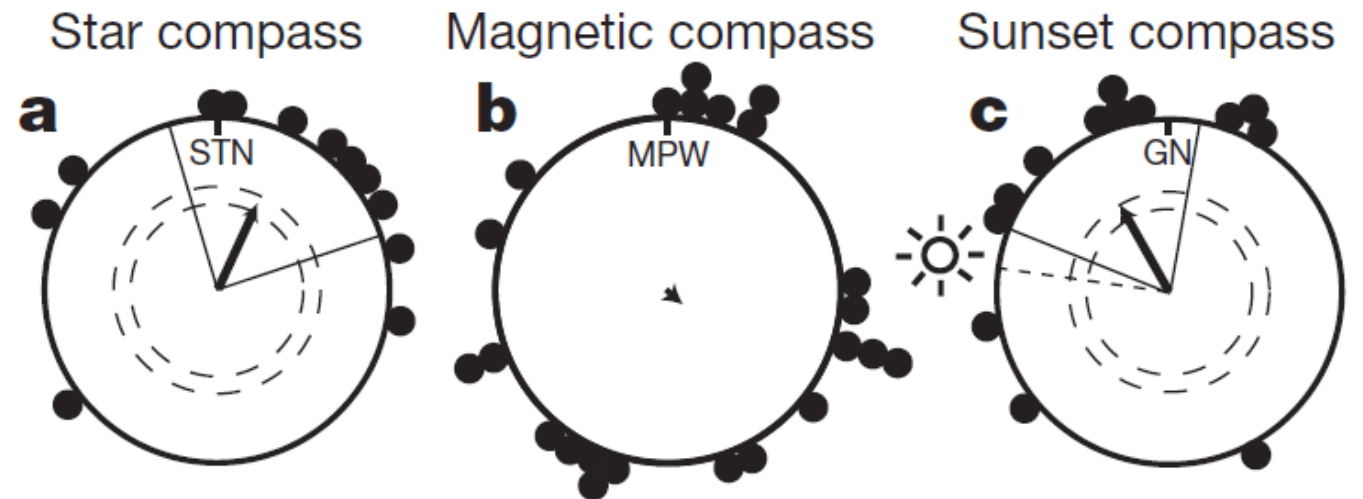
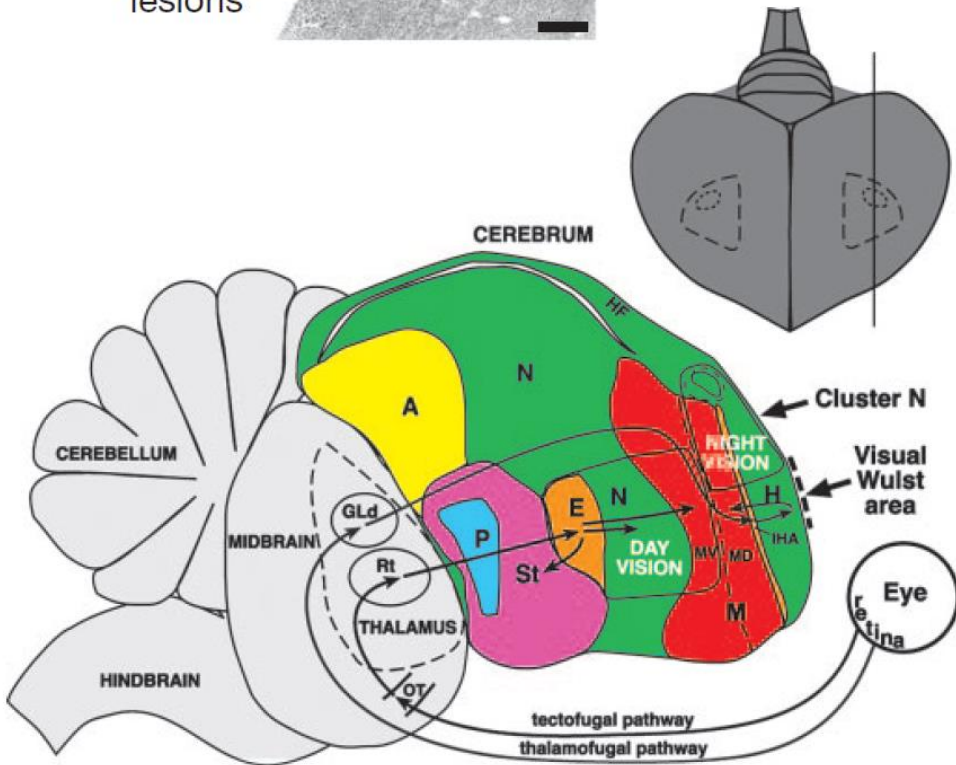
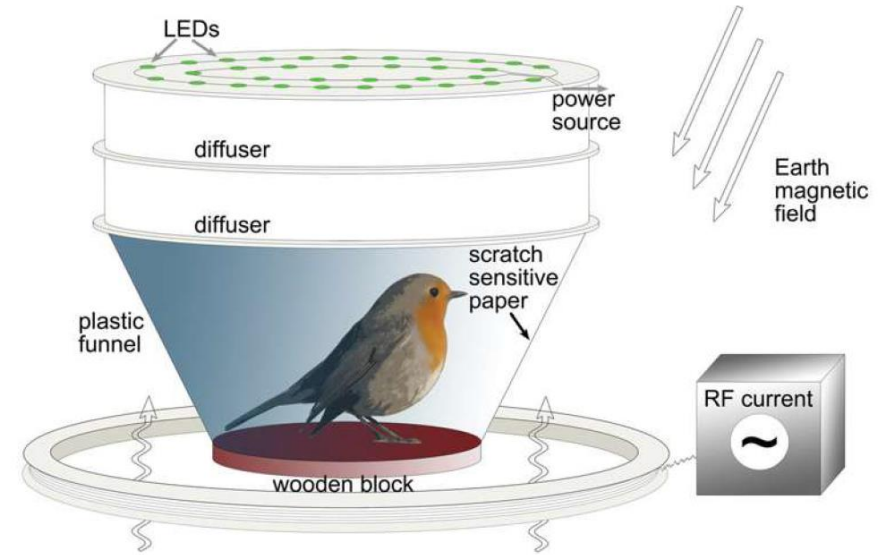
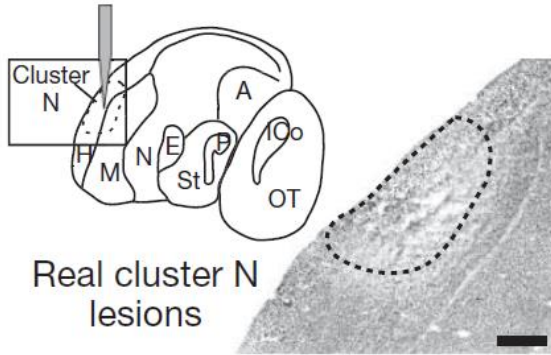




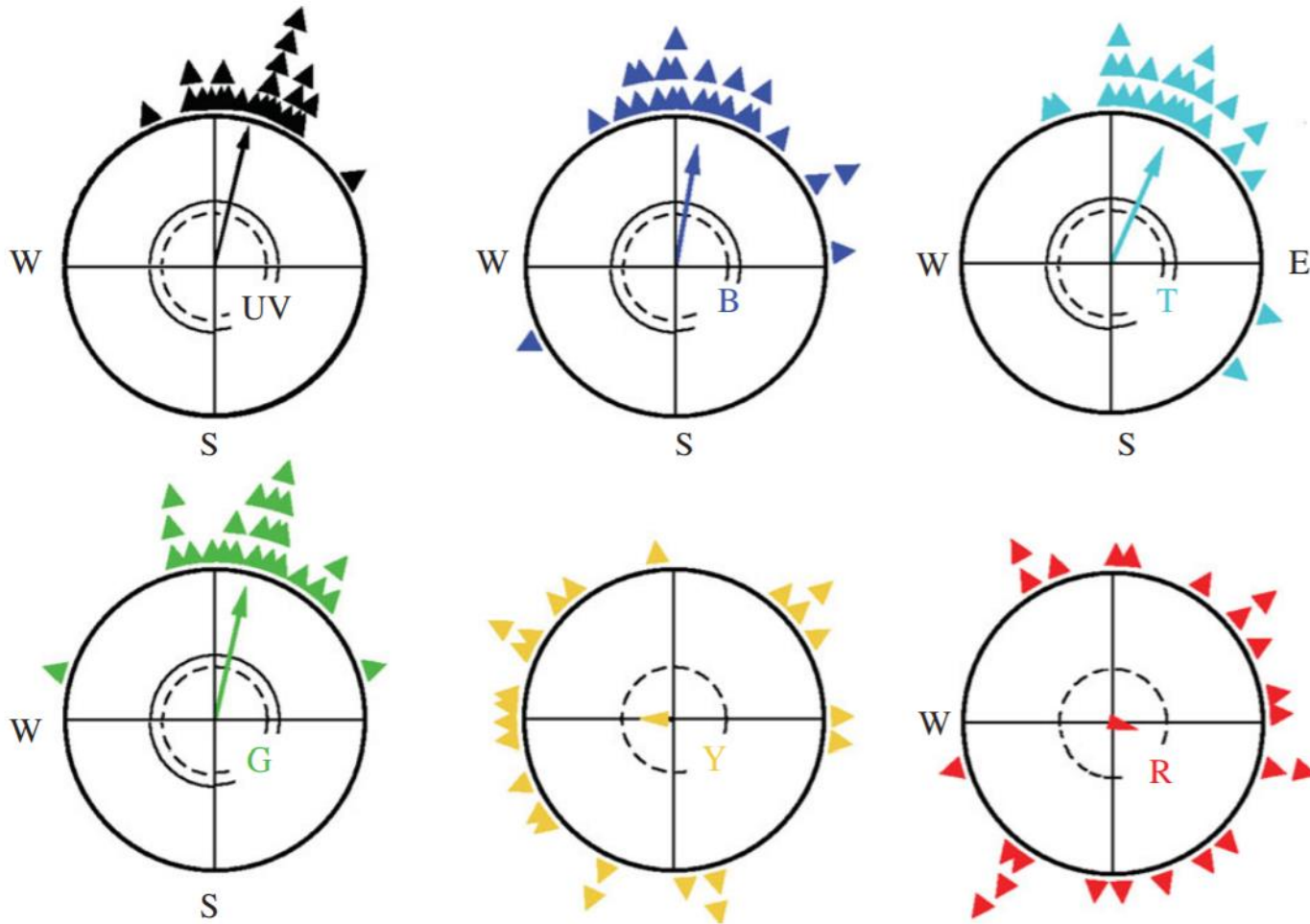
# Appendix

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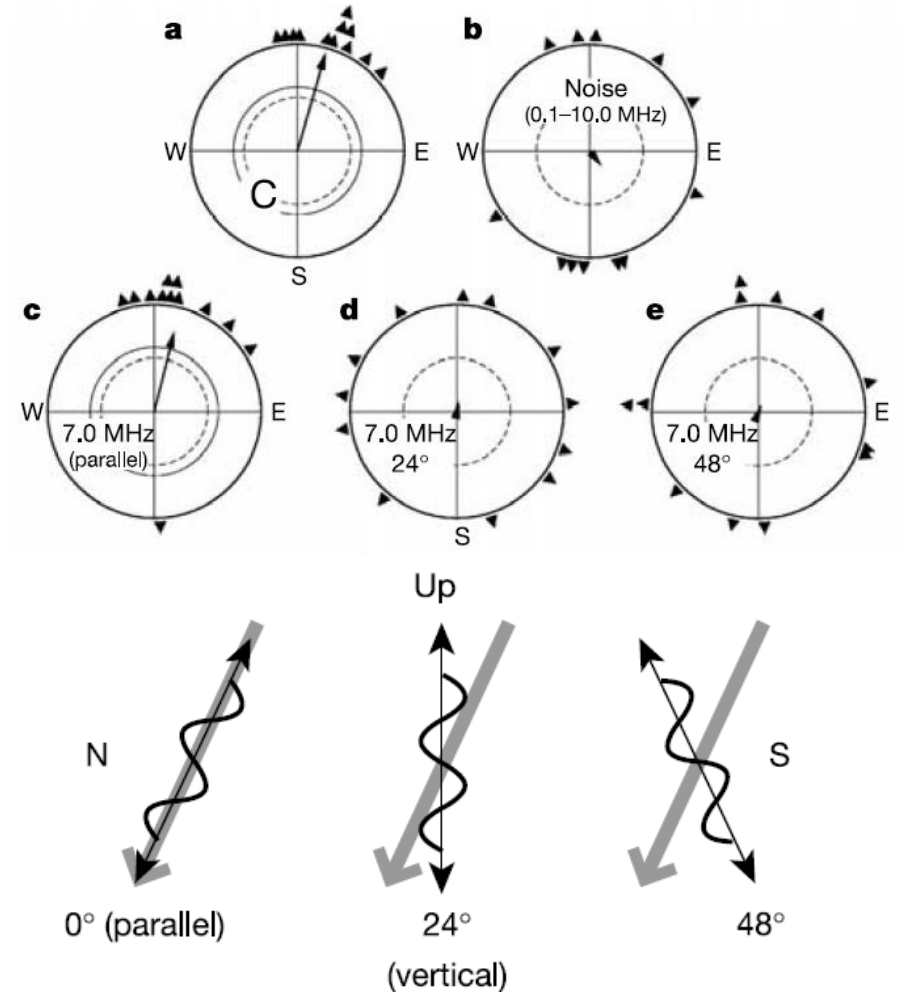
- 1) A neural connection between night vision and magnetic sensing in European robin.



- 2) Different wavelength of incident light can disrupt magnetic orientation of European robin.

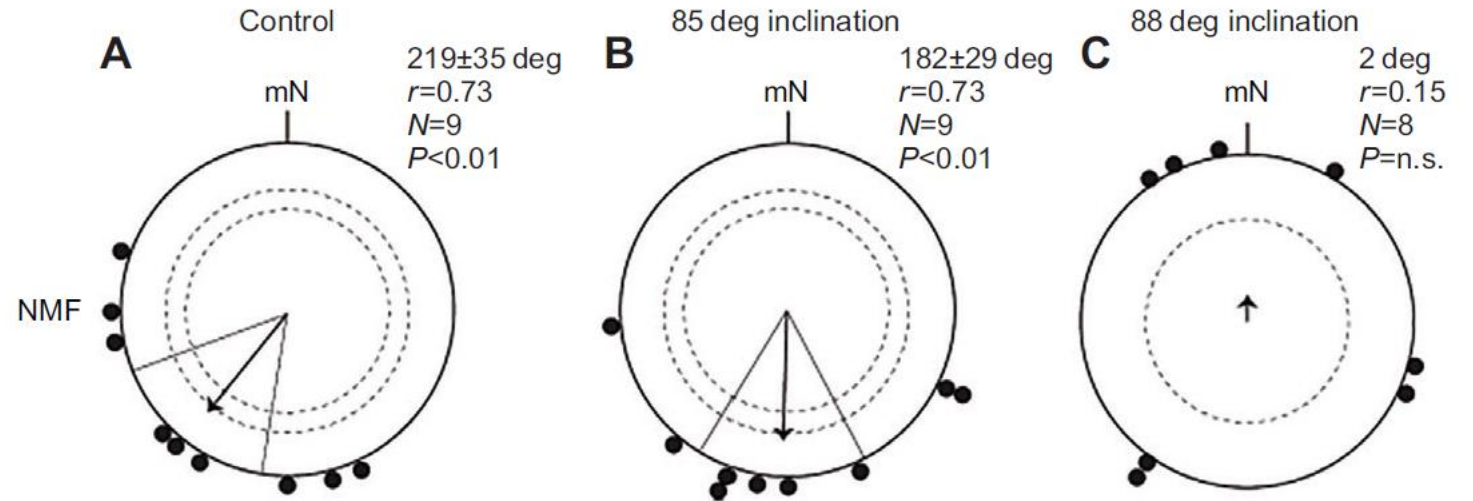
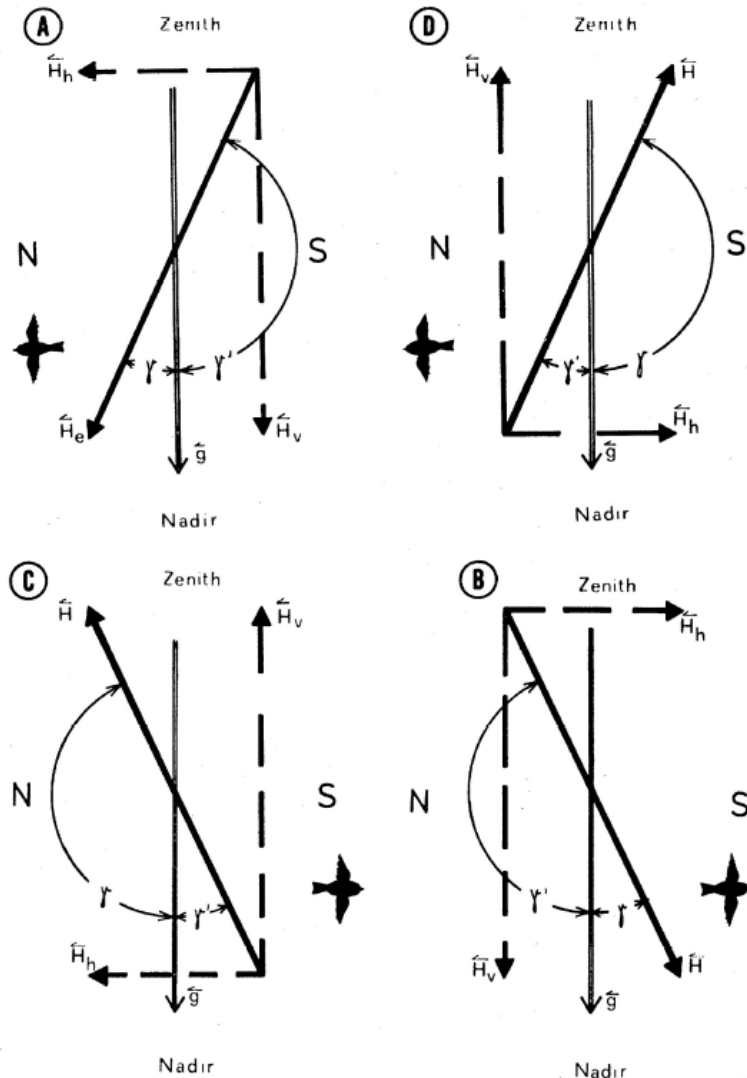


Orientation of European robin under monochromatic light of different wavelengths



Robins were disoriented when exposed to a vertically aligned broadband (0.1–10 MHz) or a single-frequency (7-MHz) field in addition to the geomagnetic field.

- 3) The magnetic compass of European robin is a inclination compass with  $<5^\circ$  precision, which can be explained by radical pair mechanism instead of any other hypothesis.



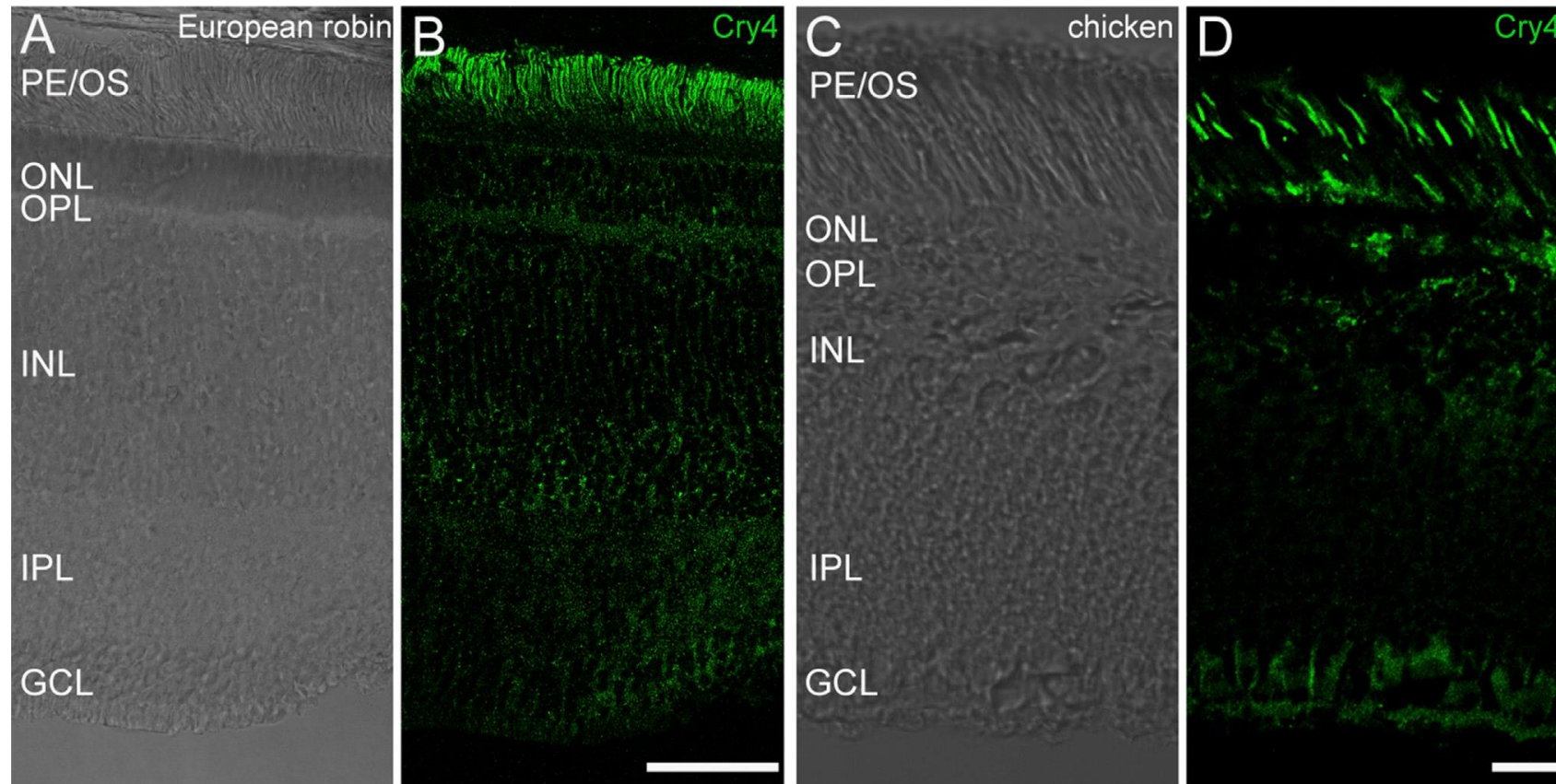
The orientation of individual blackcaps was tested in magnetic fields with 67, 85 or 88 degree inclination

The magnetic compass of European robins **does not use the polarity of magnetic field** for detecting the north direction.

The birds derive their north direction from interpreting the **inclination of the axial direction** of the magnetic field lines in space, where **field lines and gravity vector form the smaller angle**.



- 4) Cry4 protein, a member of cryptochrome family, is found recently located on the outer segment in cones in birds' retina.



**Cry4 Is Expressed in the Outer Segments of Specific Photoreceptor Cells in the Retina of European Robin and Chicken**



