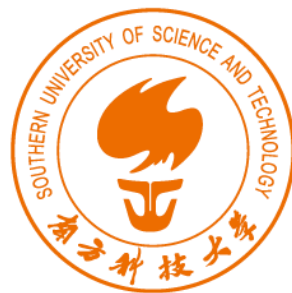


A novel biophysical model for radical pair mechanism in birds' magnetoreception



2019年5月21日

骆锦威 11410163



Contents

- Background
- Mechanisms
- OCC model
- Simulations
- Discussion

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.

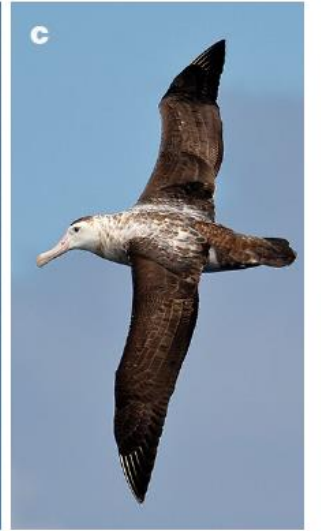
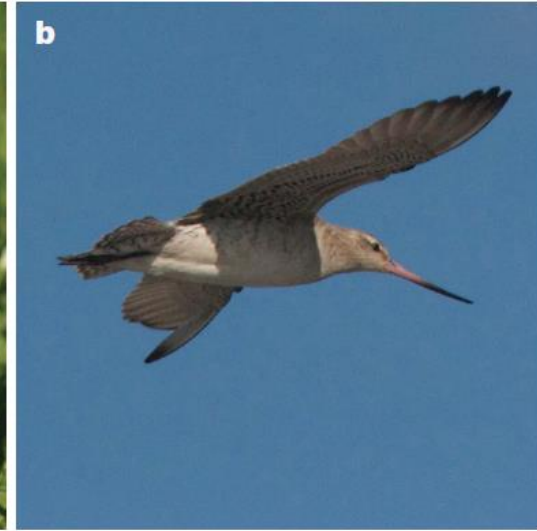
Modified after Ref.[1]

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]

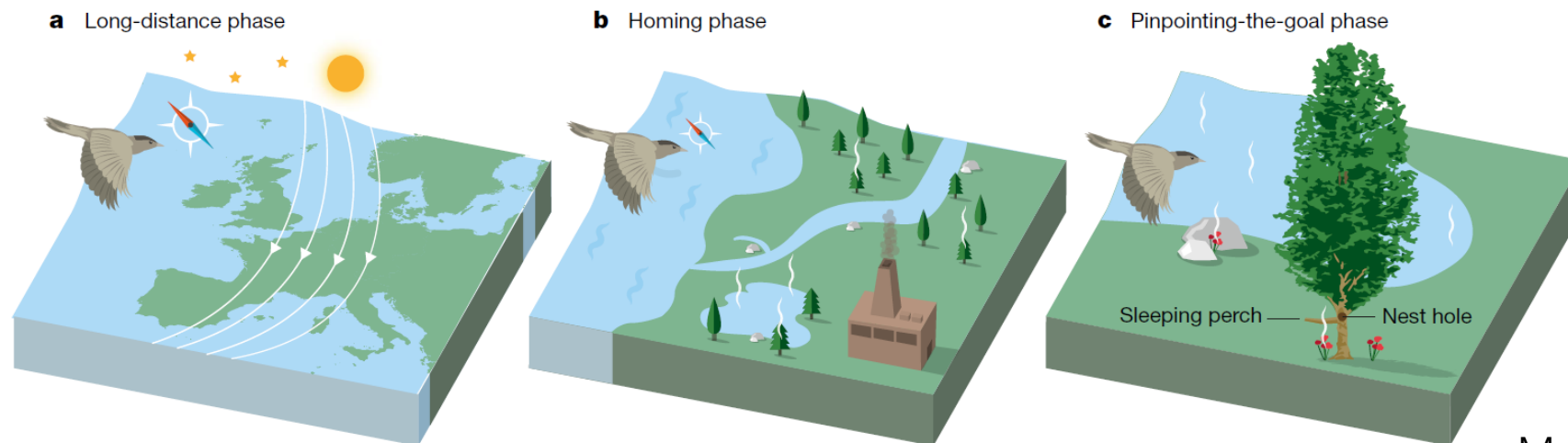
Multisensory cues for bird navigation

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

Magnetic compass, sun compass, star compass

- **Homing and pinpointing – Local orientation**

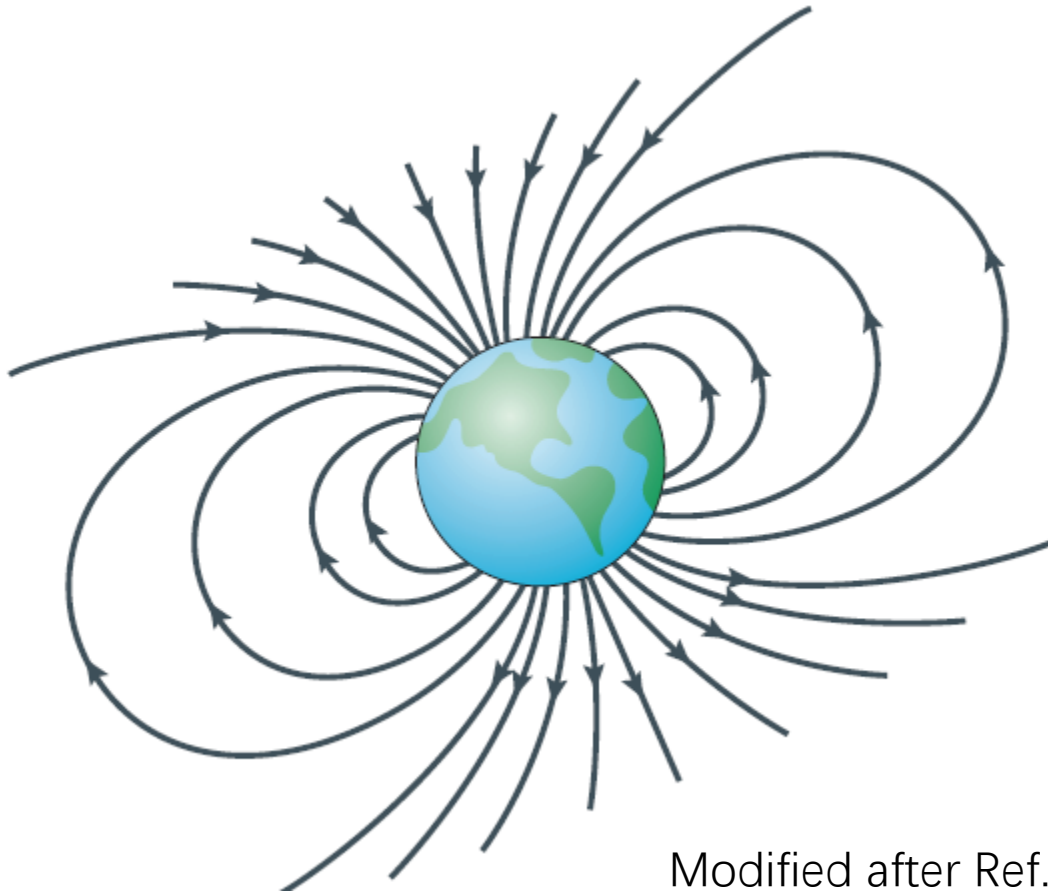
Odors, borders, coast lines, sound, landmarks such as a tree, a small hill, a specific coral



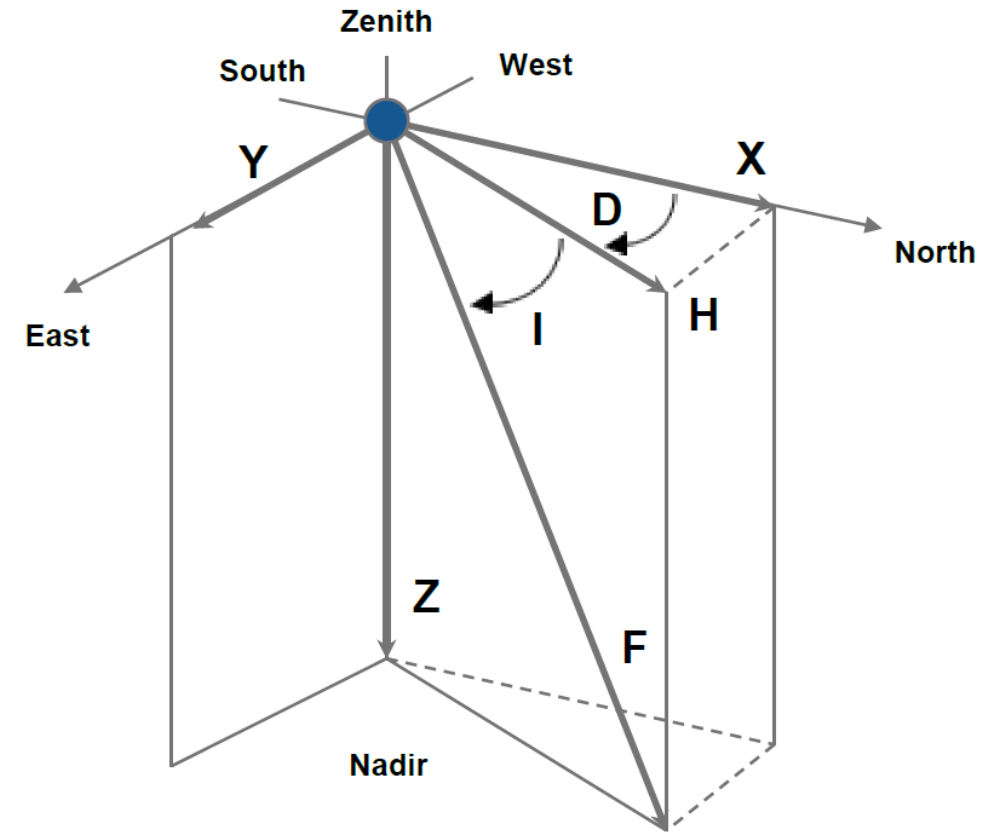
Modified after Ref.[2]

Earth magnetic fields

- Field strength: 25~60uT



Modified after Ref.[3]



Magnetic field information

Inclination

Polarity/Declination

Intensity

Mechanisms

Three possible mechanisms

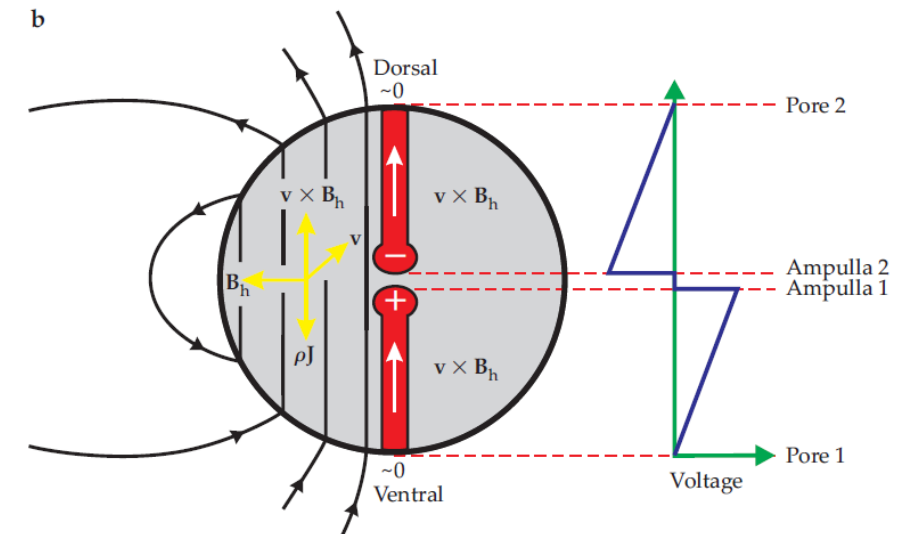
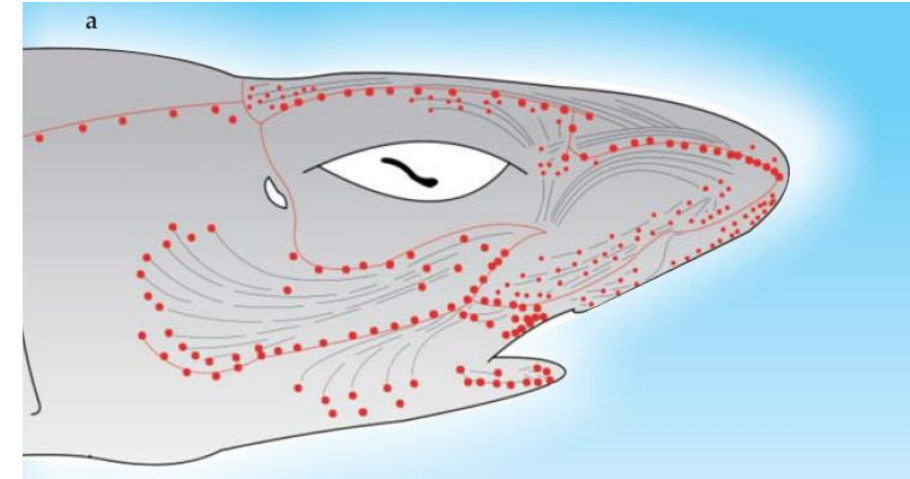
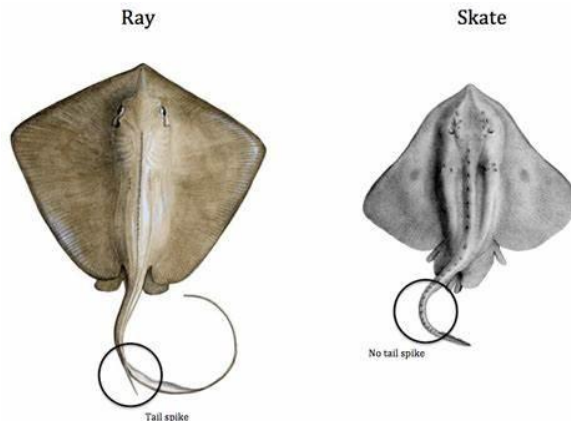
- Electromagnetic induction
- Magnetite based magnetoreception
- Radical based magnetoreception

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

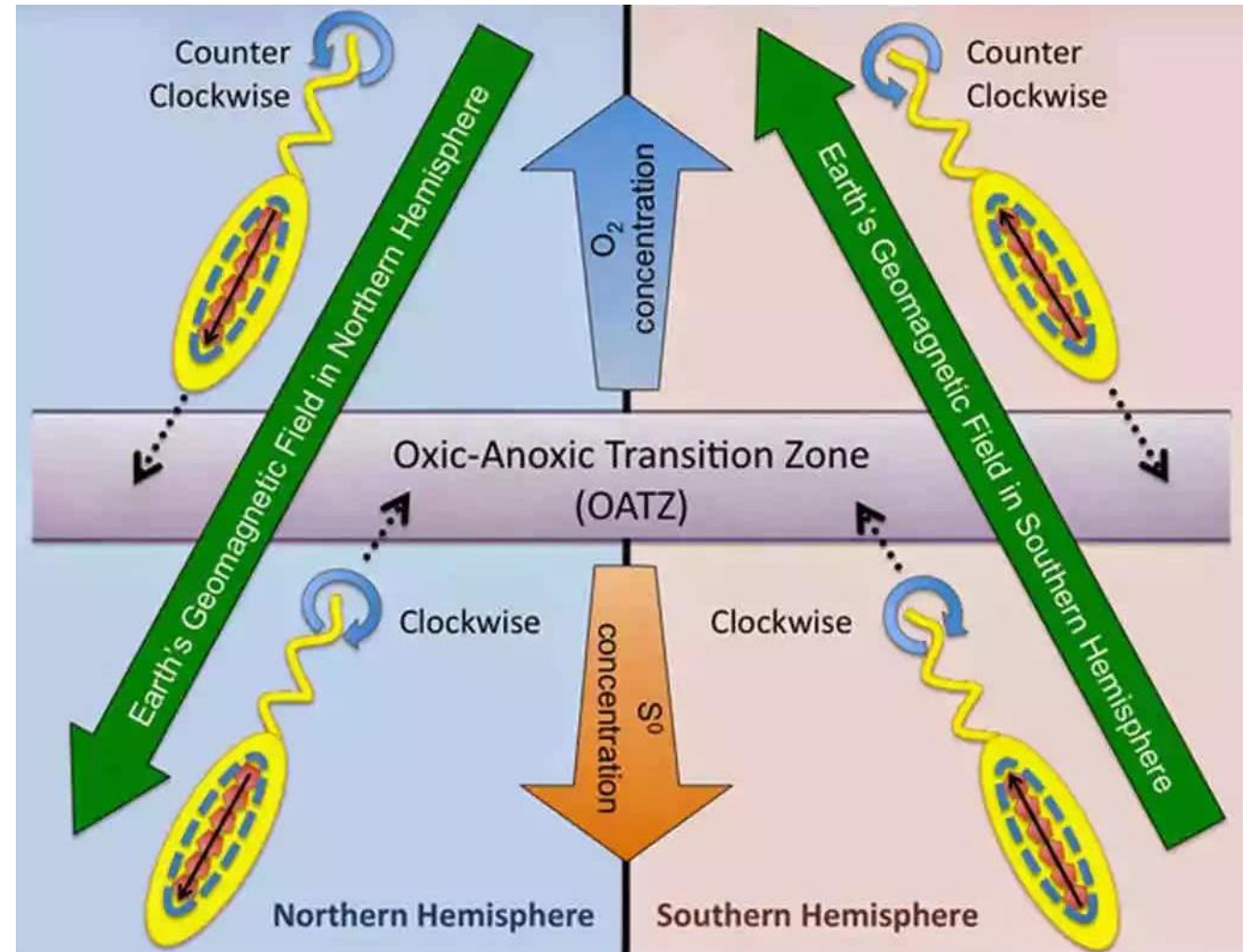
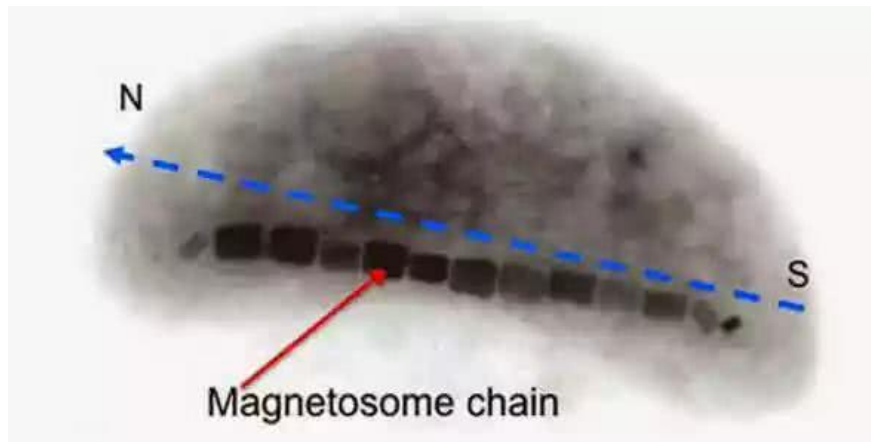
- A unique organ in cartilaginous fish:
- **Ampullae of Lorenzini**
- Jelly-filled pores with electroreceptor around
- Moving magnetic field accumulates electric potential on two sides of a pore via proton flux, which can be detected by electroreceptor cells.
- Elasmobranch species: sharks, rays and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



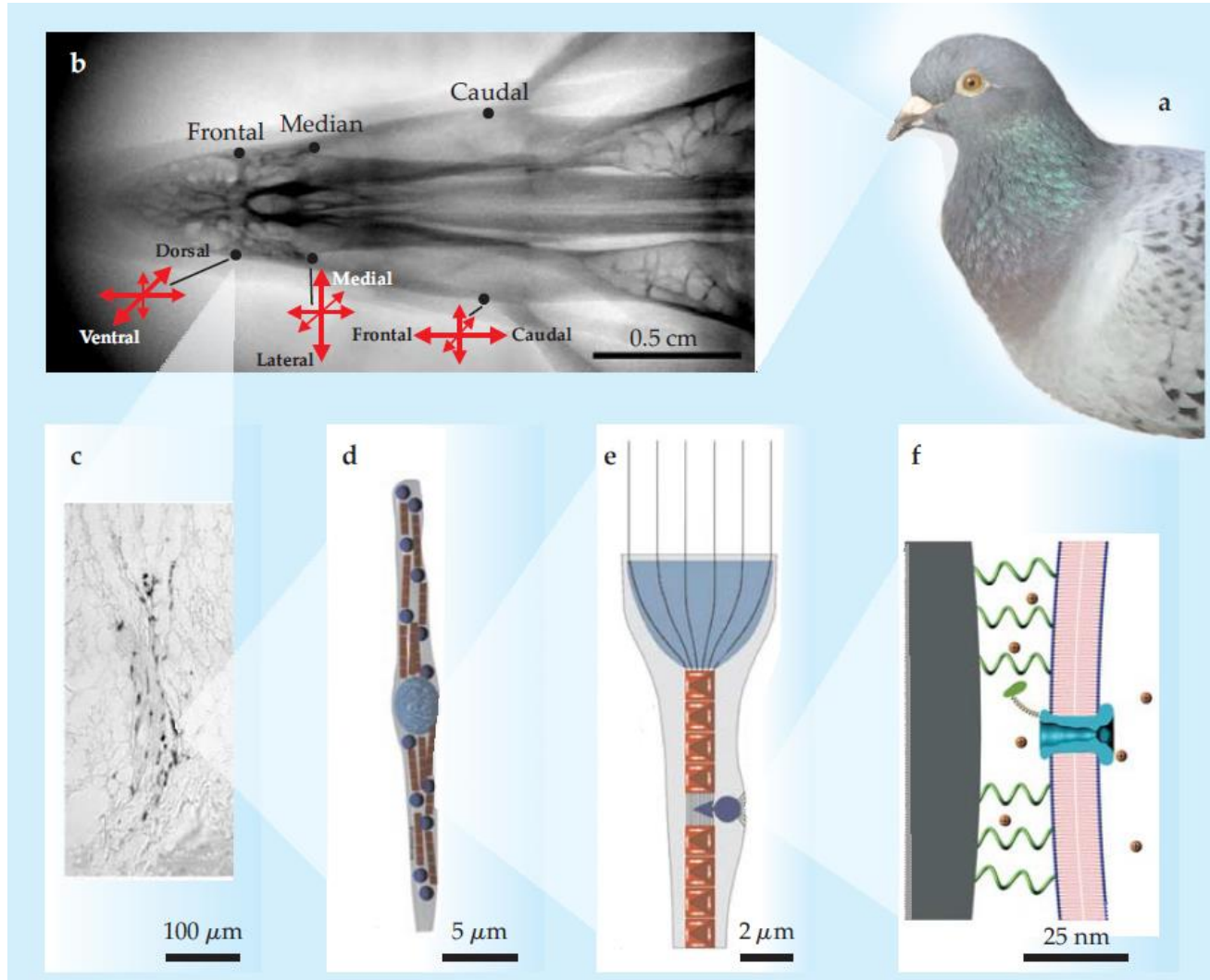
Modified after Ref.[4]

Magnetite based Magnetoreception

- Magnetotactic bacteria
- A magnetosome chain:
- 15-20 Fe_3S_4 , Fe_3O_4 crystal
- 30-100nm for each crystal

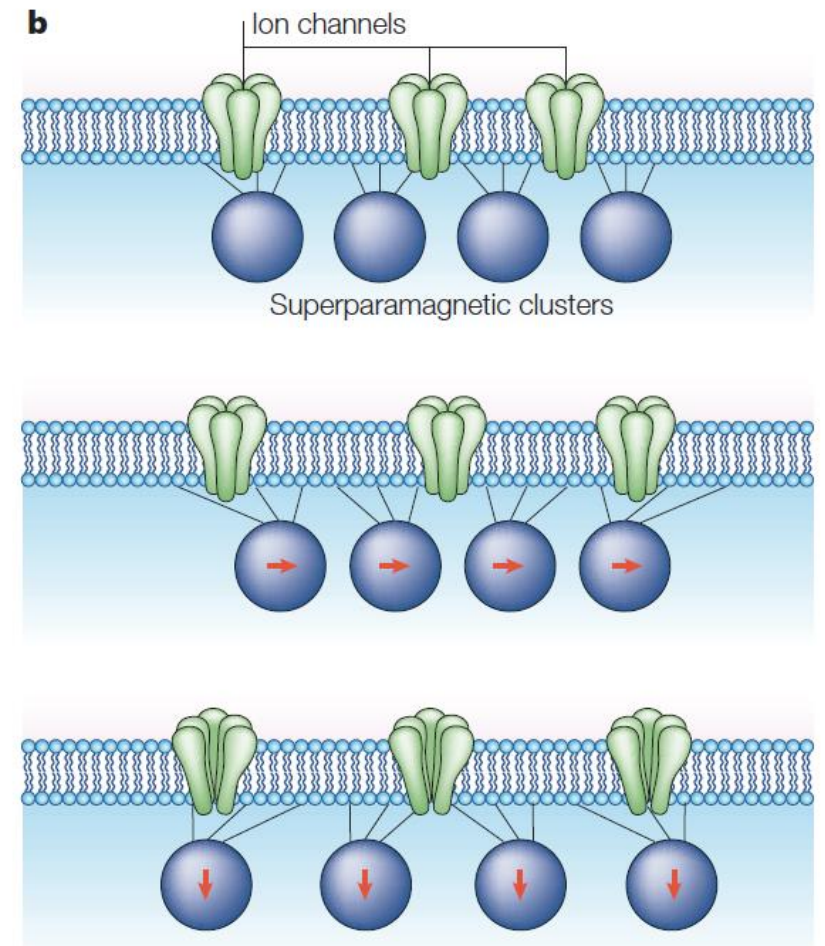


Magnetite based Magnetoreception



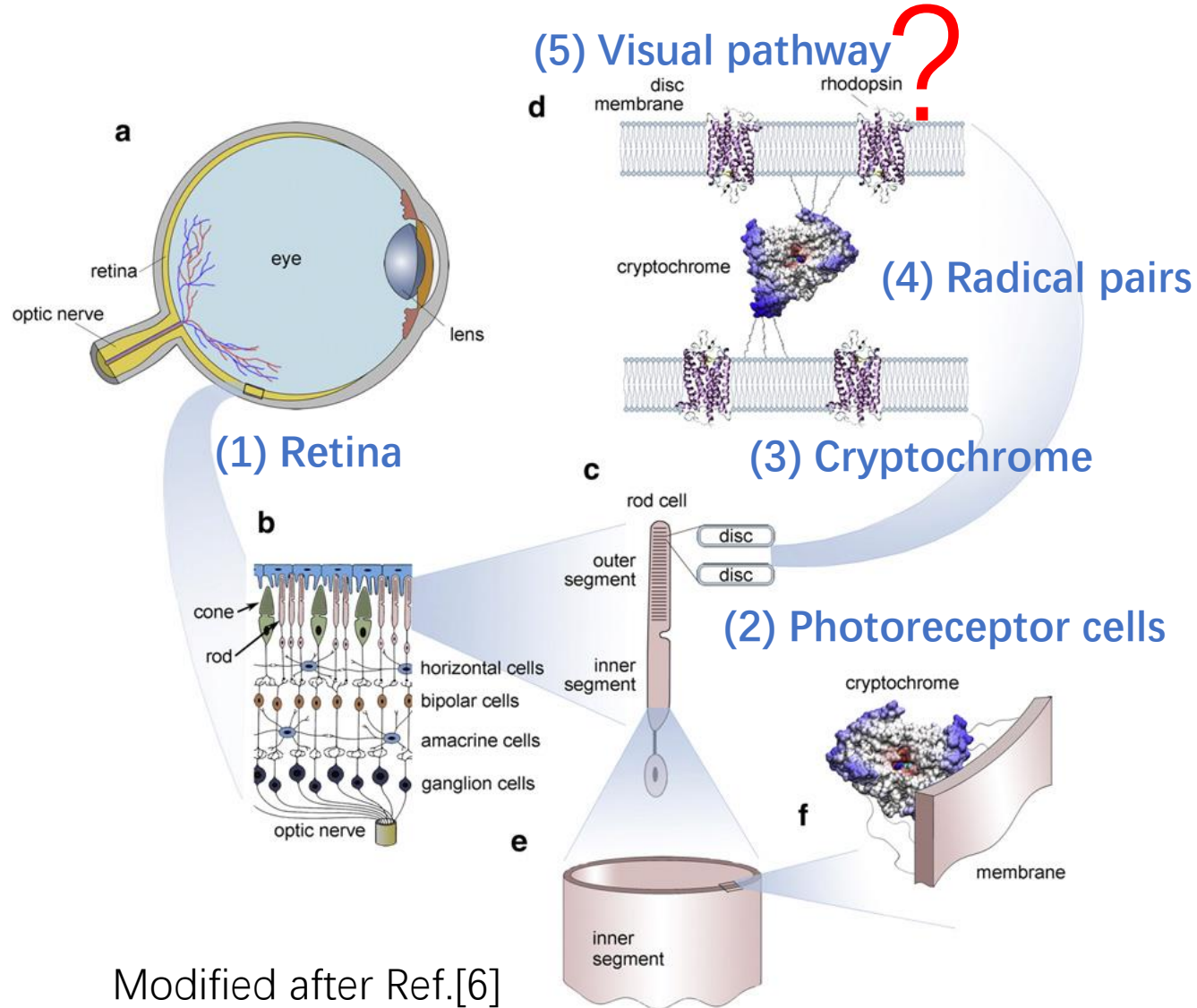
Modified after Ref.[4]

Superparamagnetic crystals



Modified after Ref.[3]

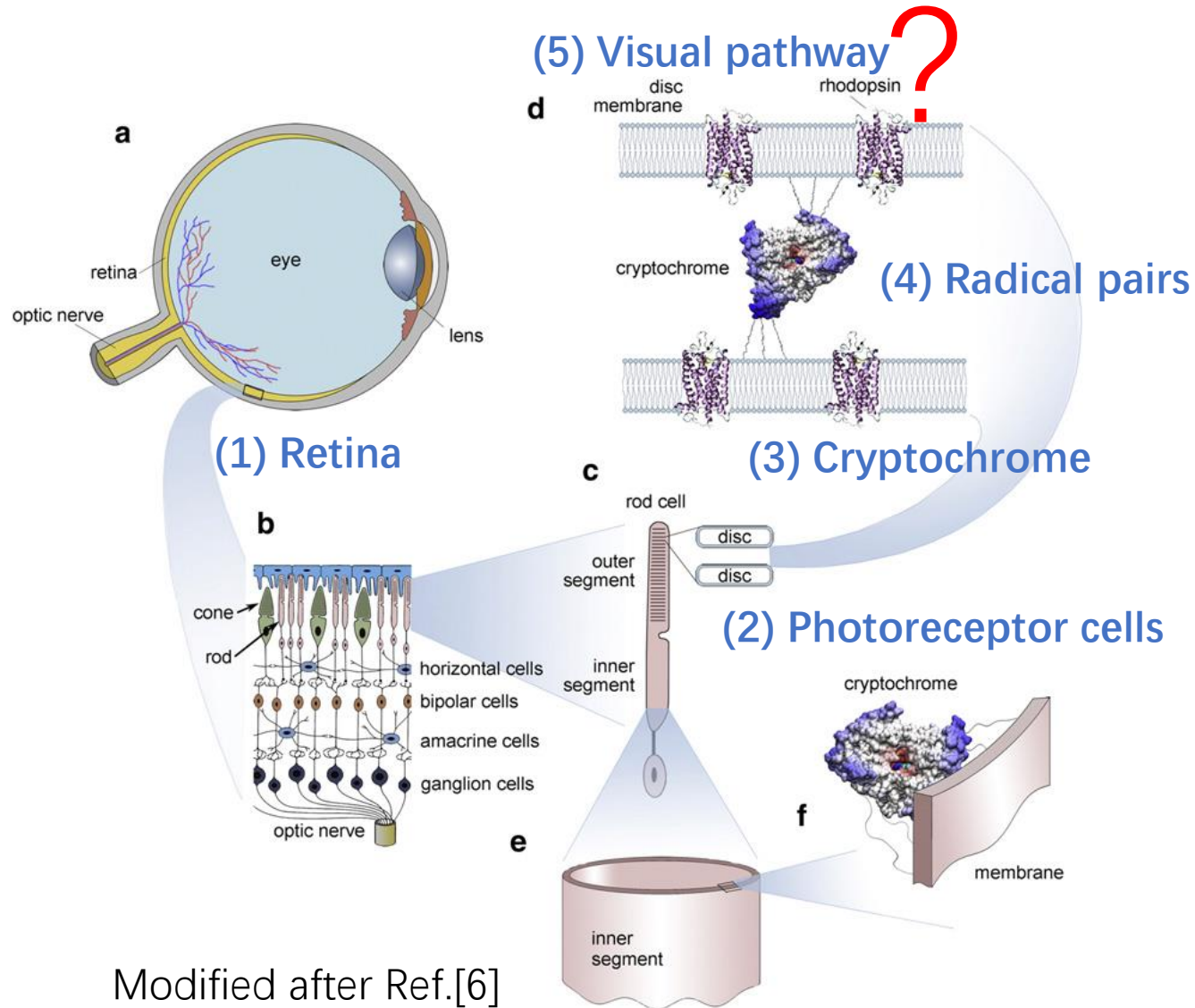
Radical based magnetoreception



Modified after Ref.[6]

- 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.
- 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.
- 4) Cryptochrome is the **only protein family** that can form **a light-induced radical pair**.

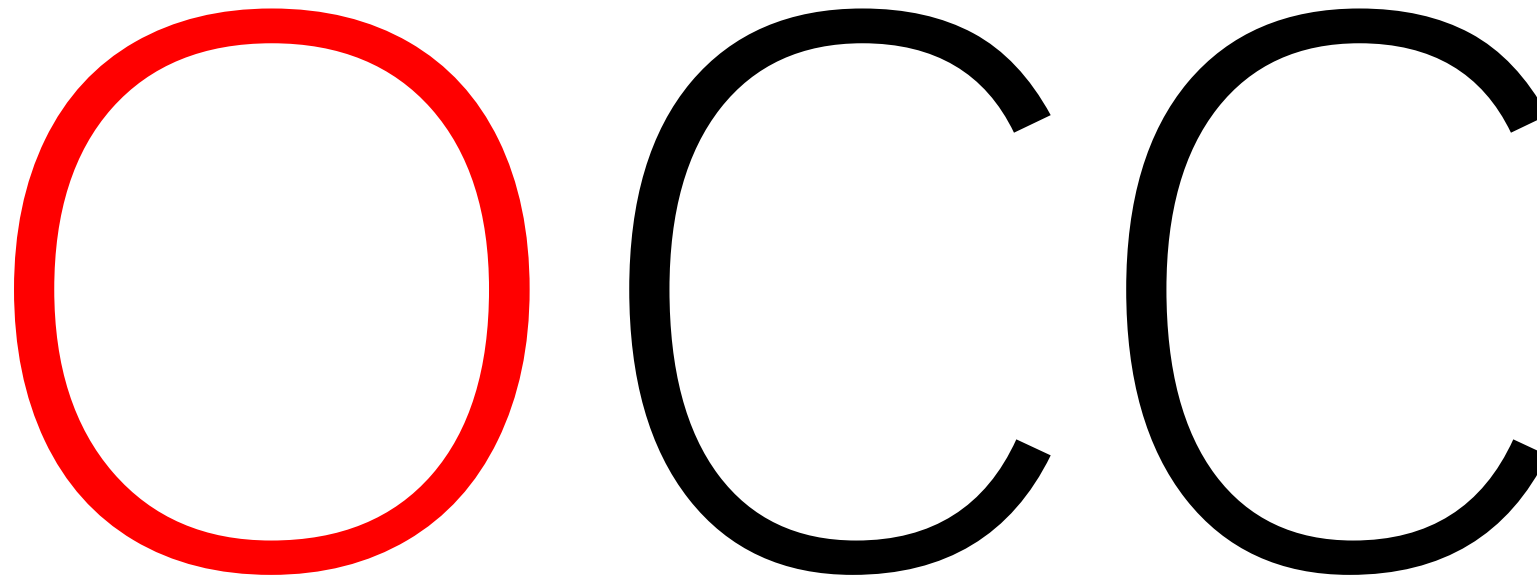
Radical based magnetoreception



Modified after Ref.[6]

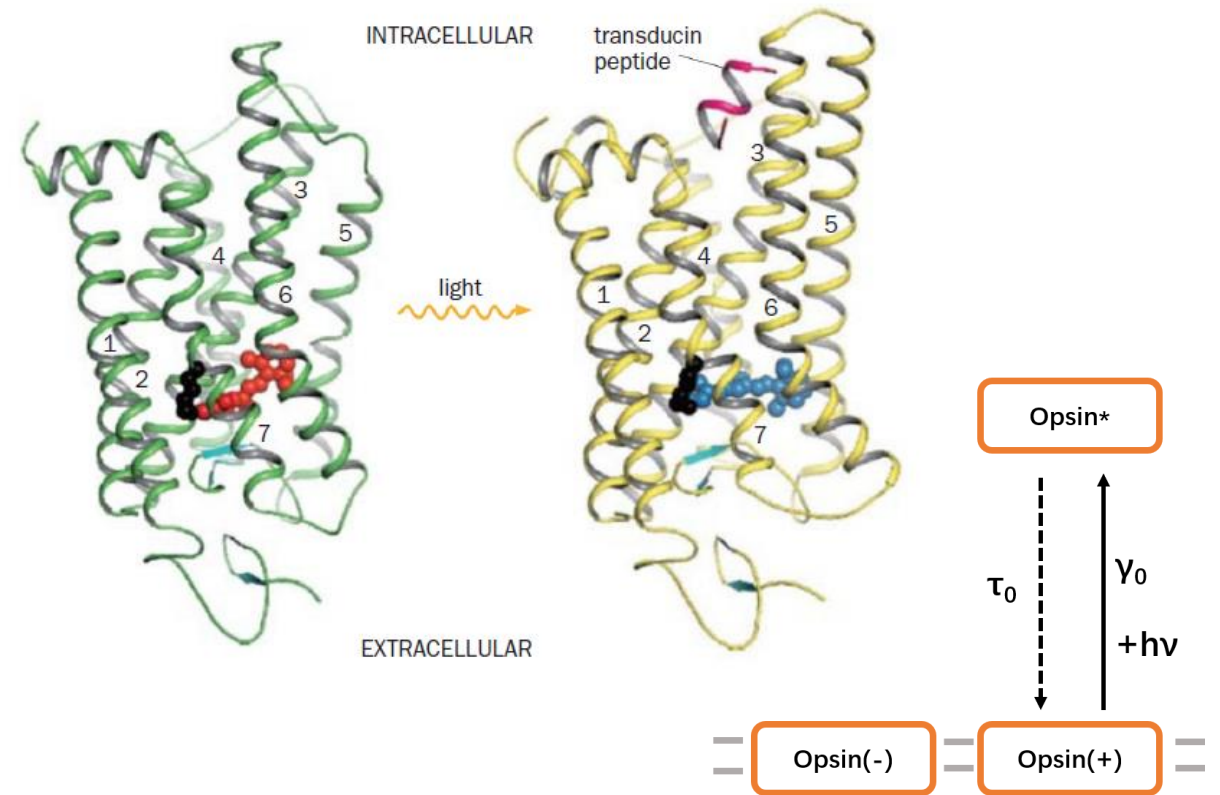
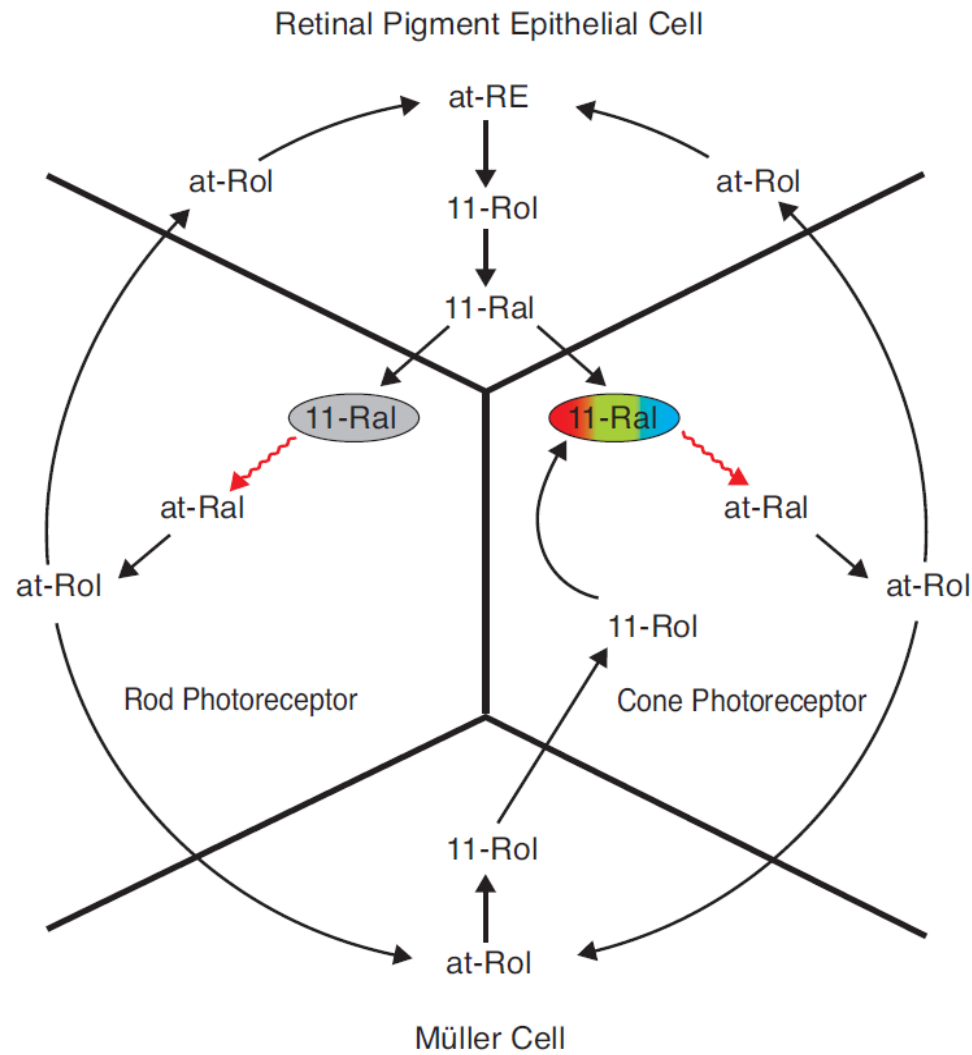
- 5) **Cry4 protein**, a member of cryptochrome family, is found recently located on the **outer segment** in cones in birds' retina.
- 6) A popular hypothesis of (4)-(5) is an noncanonical biological pathway of visual system that can modulate night vision. **No such signaling pathway** is found till 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

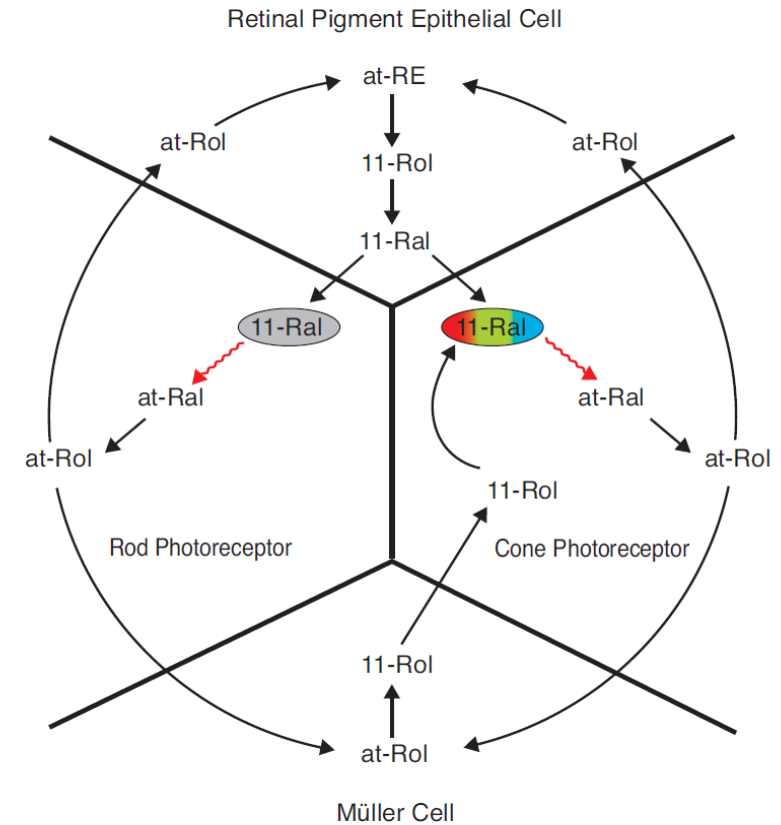
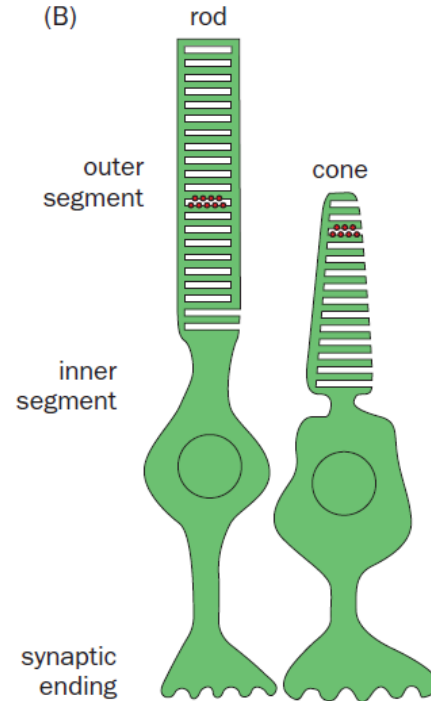
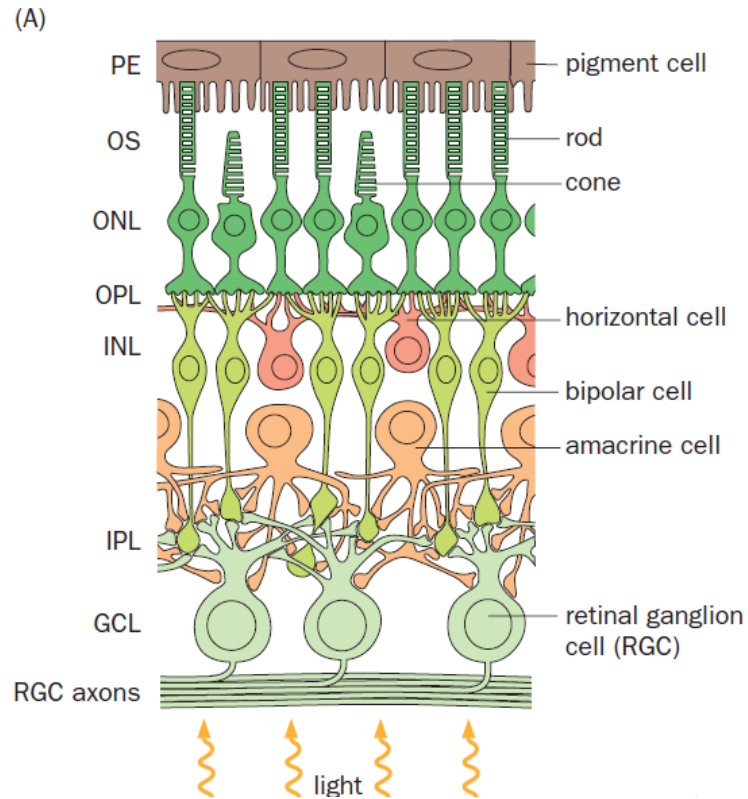
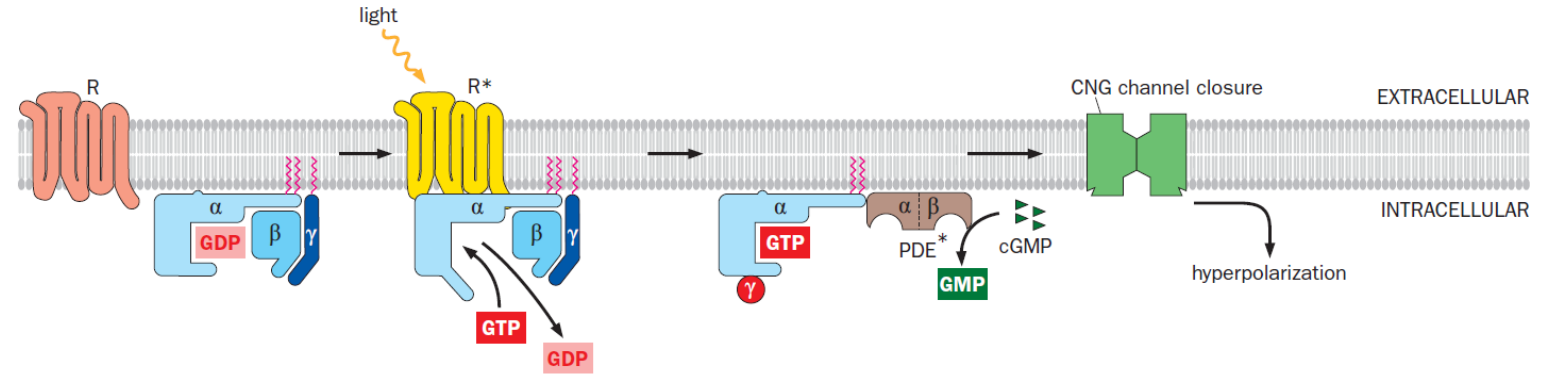
A rod-visual cycle: cis-retinal \leftrightarrow trans-retinal



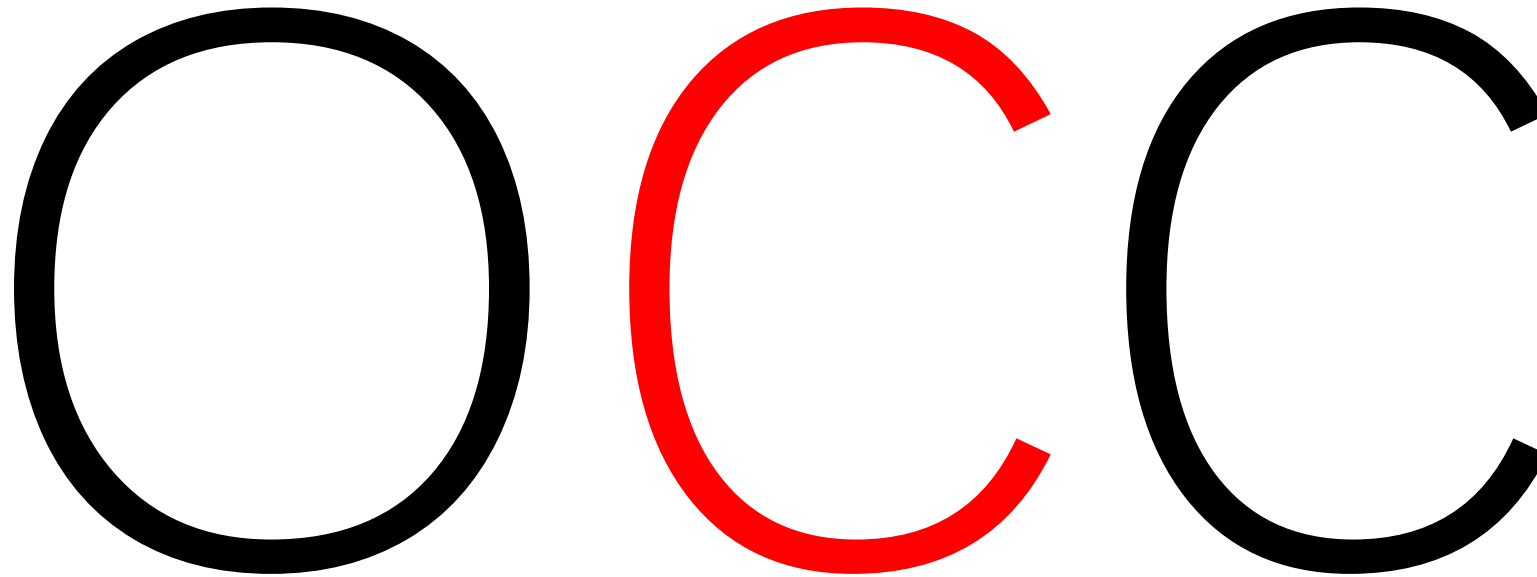
Abbreviations: at-RE, all-*trans*-retinyl esters; at-Ral, all-*trans*-retinal; at-Rol, all-*trans*-retinol; 11-Ral, 11-*cis*-retinal; 11-Rol, 11-*cis*-retinol.

Modified after Ref.[7,8]

Light perception

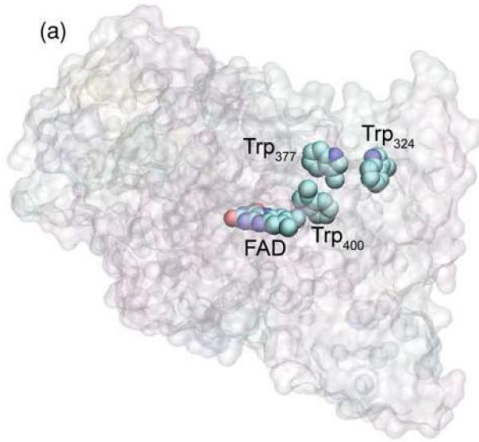


Modified after Ref.[7,8]

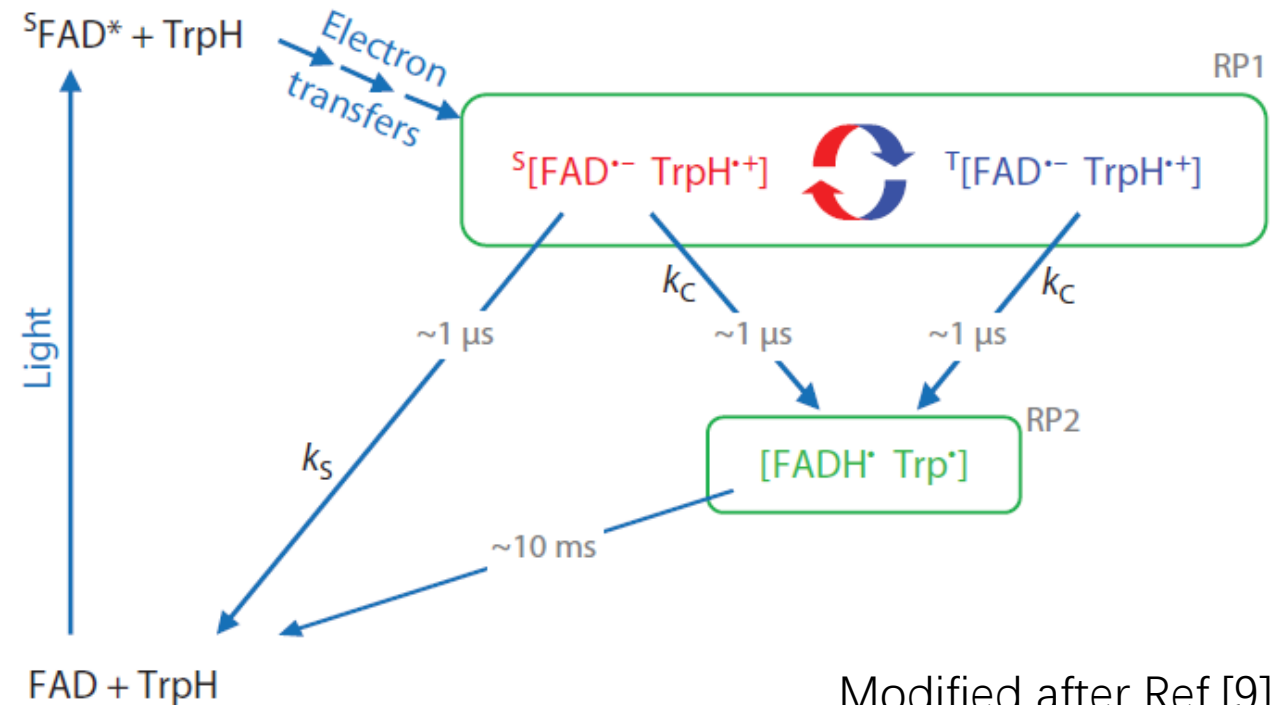
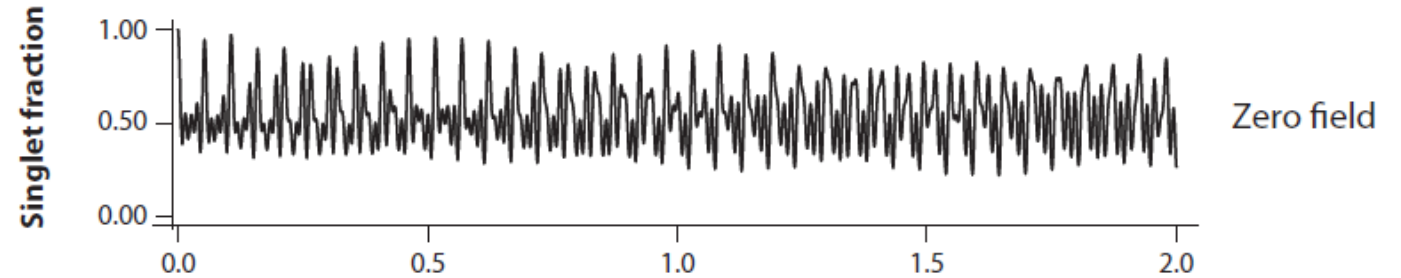
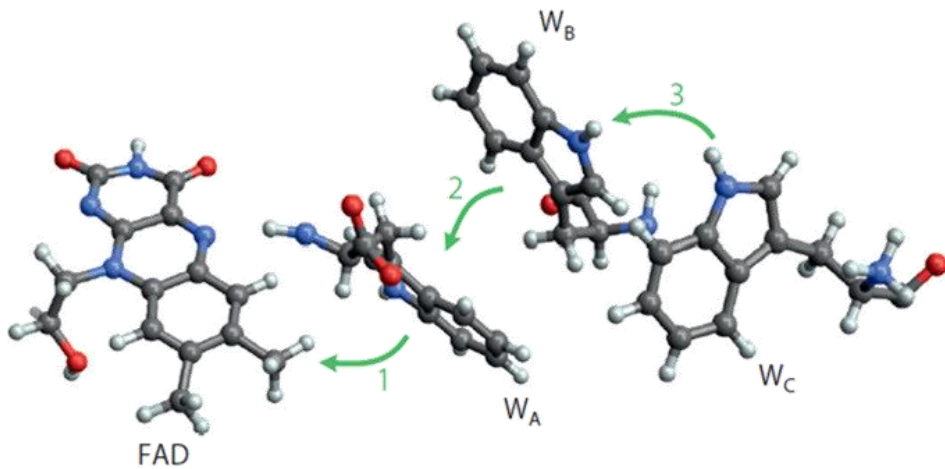


Cryptochrome, Radical pairs

Principle of radical pair mechanism

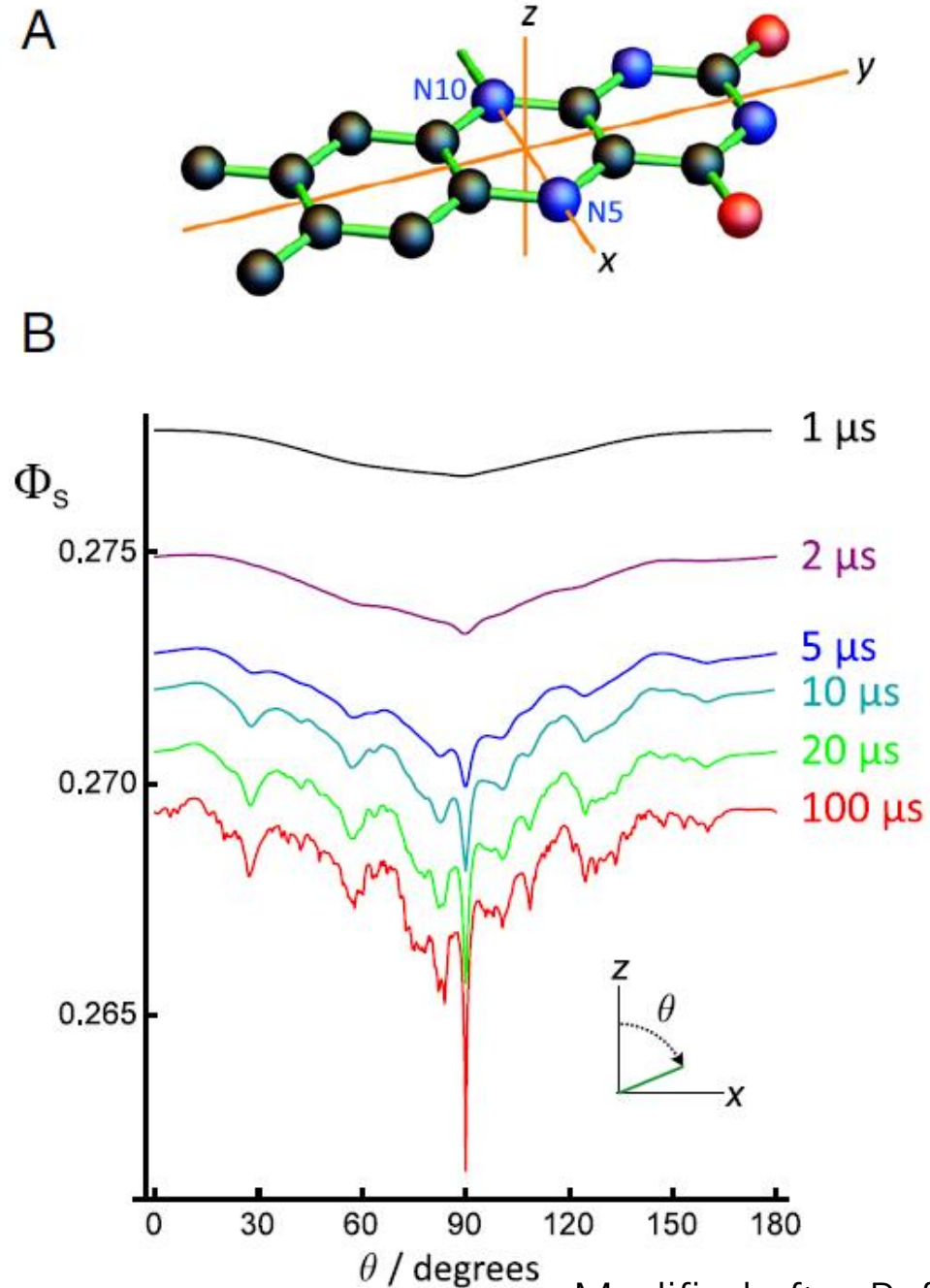
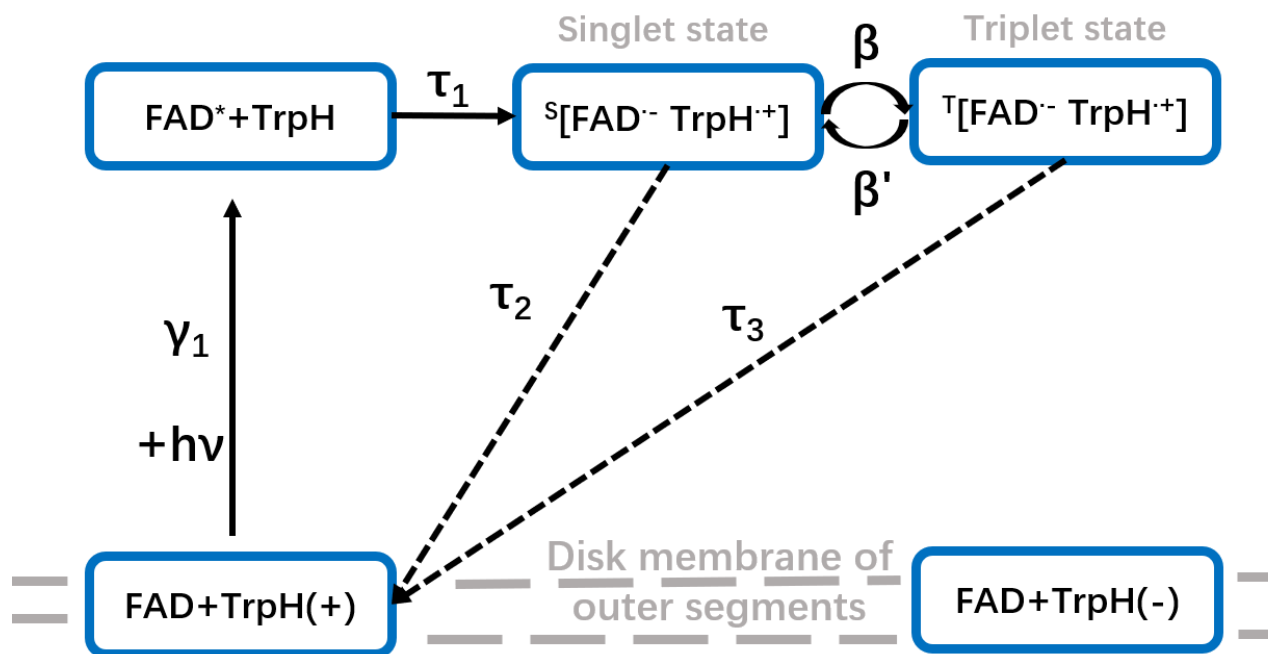
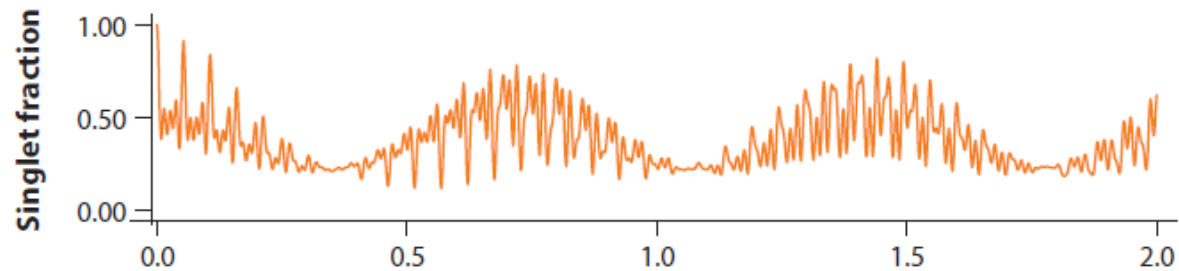


Cryptochrome



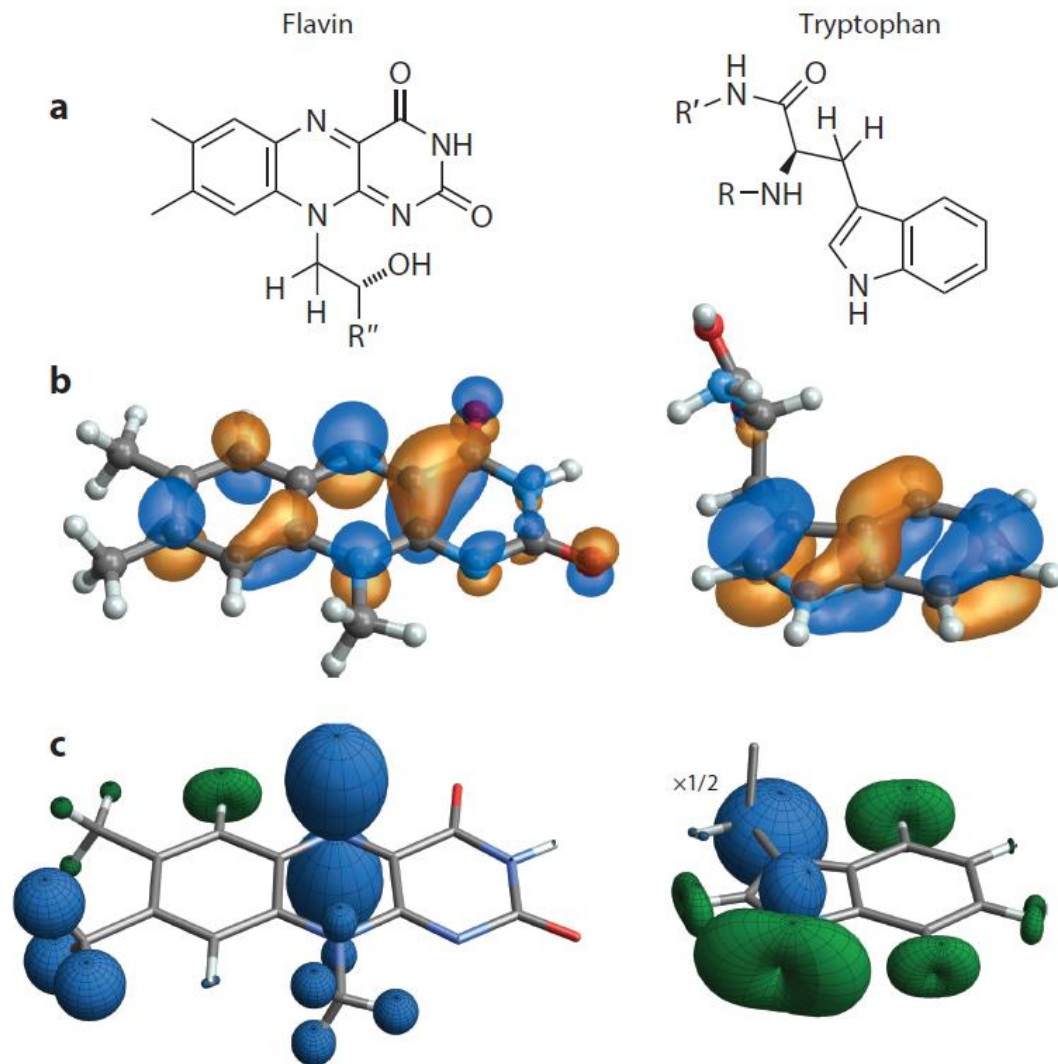
Modified after Ref.[9]

Spin dynamics



Modified after Ref.[10]

Quantum effect



Modified after Ref.[10]

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.

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

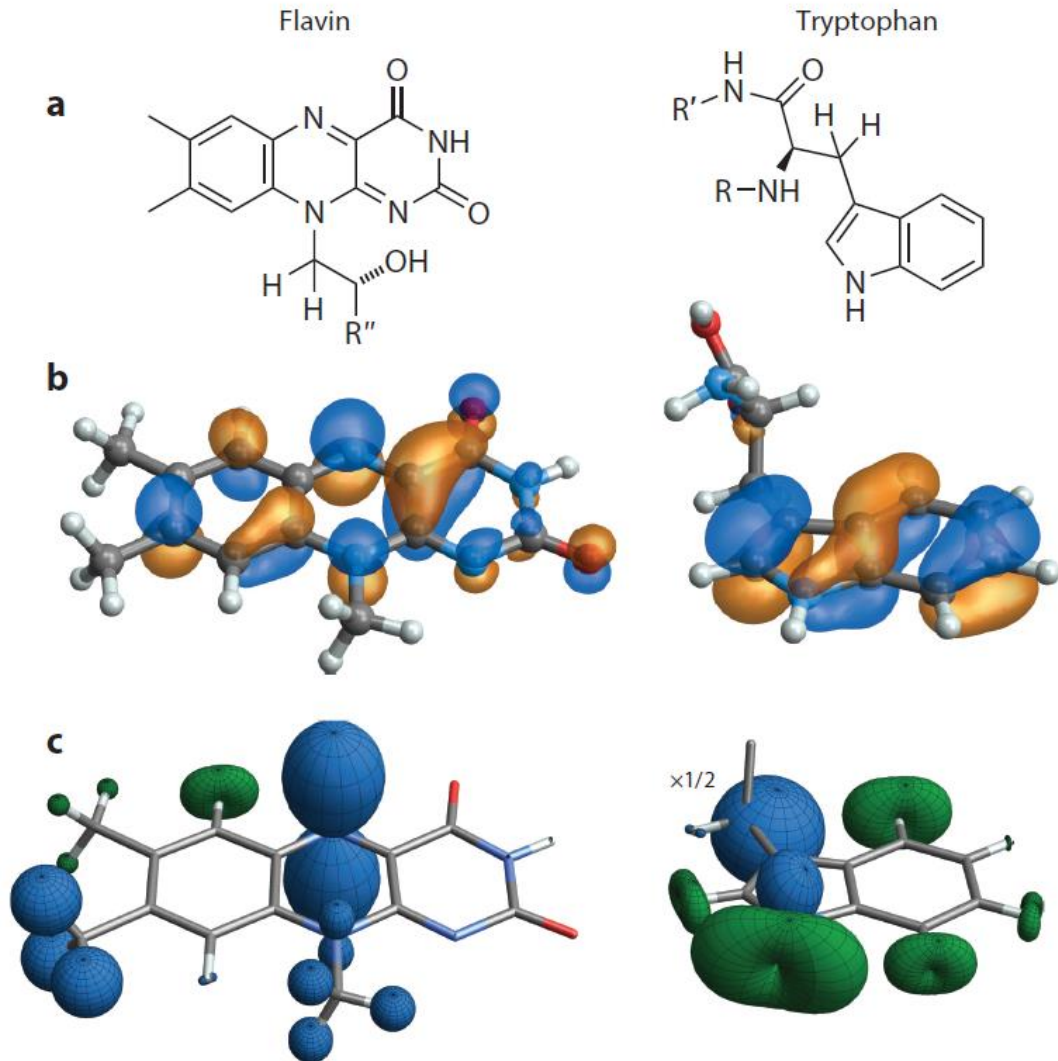
$$H_j = g\mu_B \vec{S}_j \cdot (\vec{B} + A_j \vec{I}_j)$$

\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.

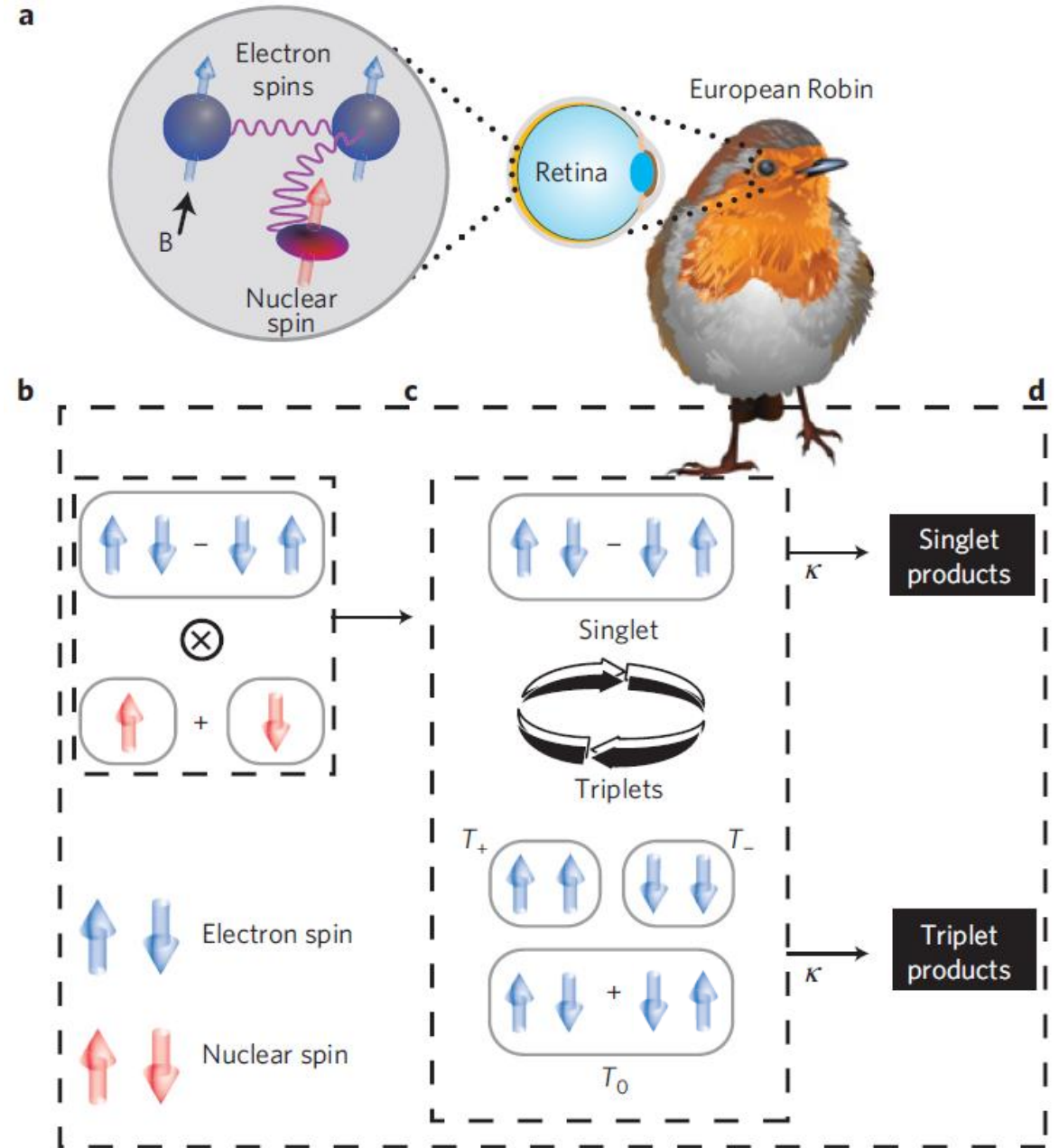
$$T(t) = \phi_S = 1 - \text{Tr}[Q^T \frac{1}{N} e^{-\frac{iHt}{\hbar}} Q^S e^{-kt} e^{\frac{iHt}{\hbar}}]$$

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

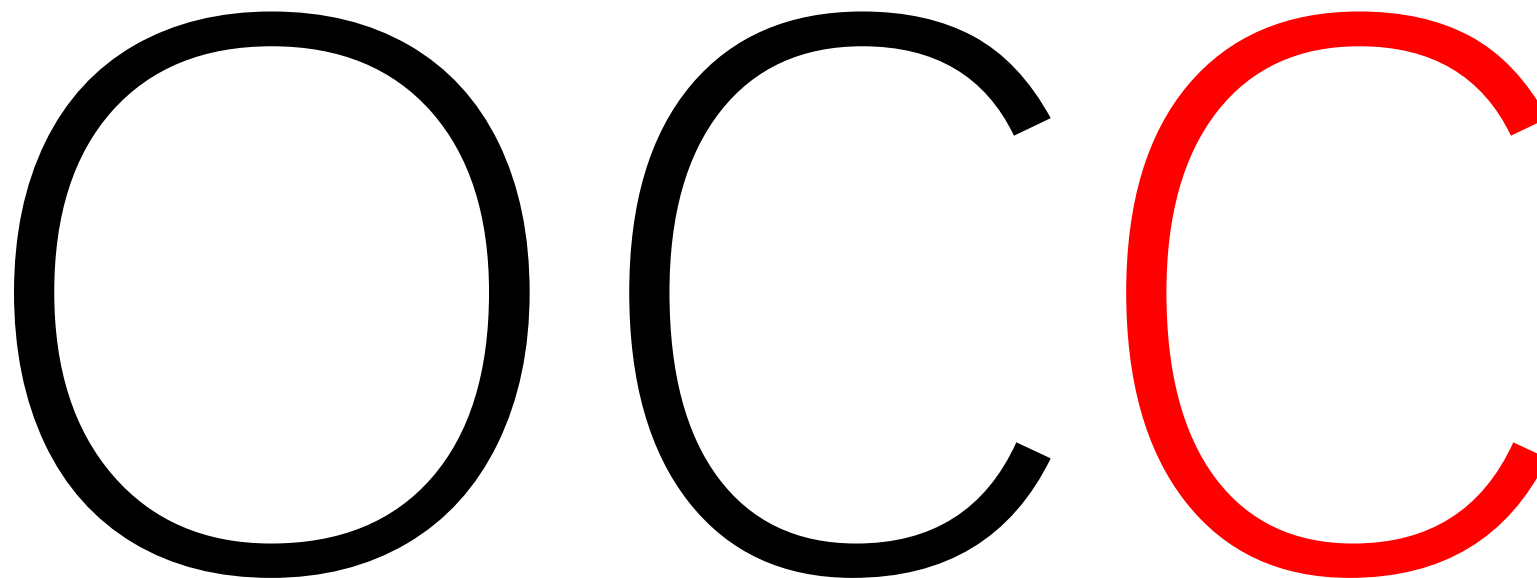
Quantum effect



Modified after Ref.[10]

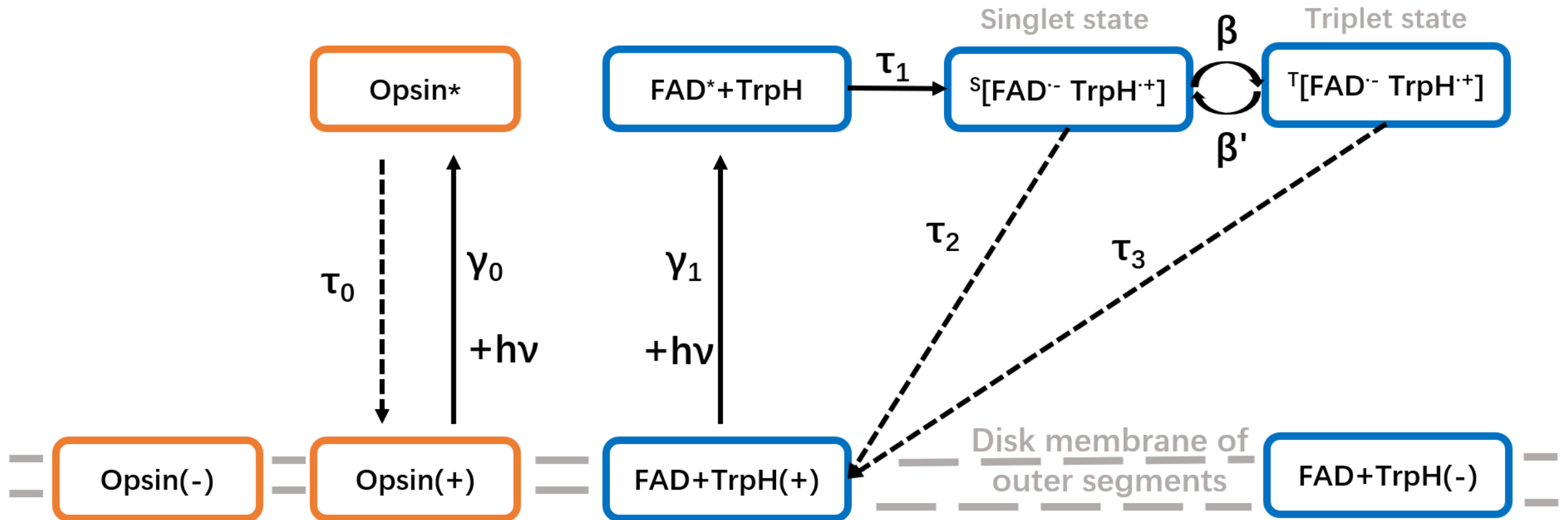


Modified after Ref.[11]



Photon Competition

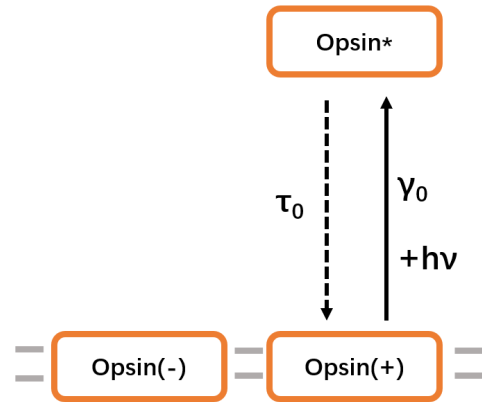
Opsin-Cryptochrome Competition model



Model formulation

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

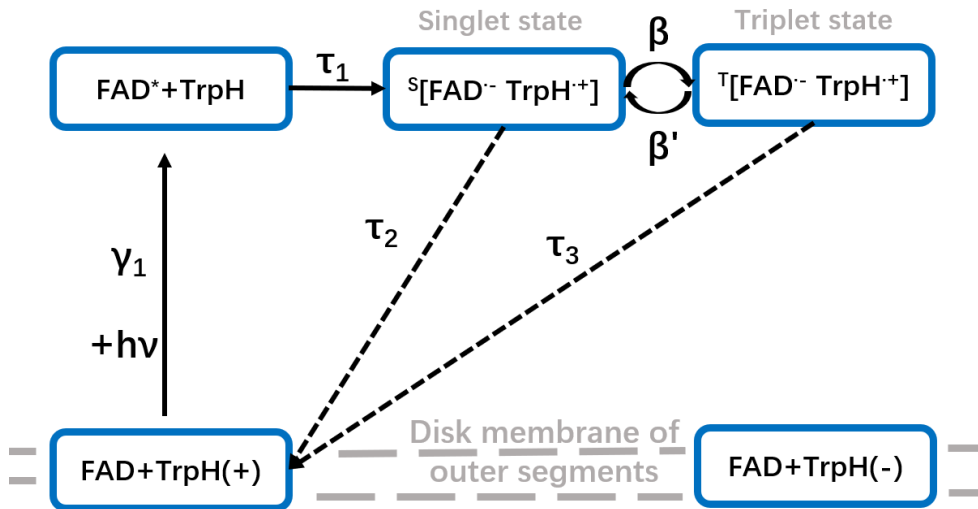
$$R_{opsin} = \frac{N_{op}^{(+)}}{N_{photon}} \quad (10)$$



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

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

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



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

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

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

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

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

A stationary solution: let all derivatives=0



Simulation

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

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

$$B = \tau_0 \gamma_0 + 1 \quad (13)$$

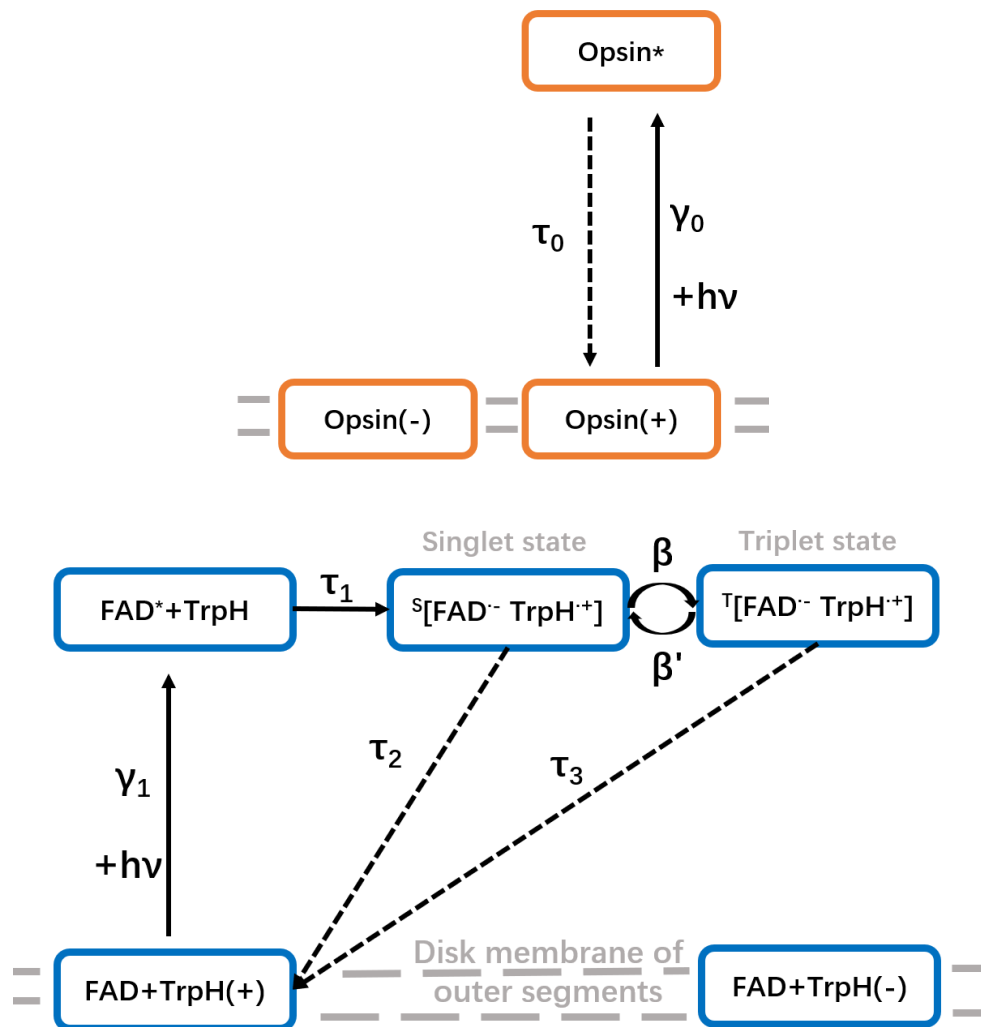
$$A = 1 + \gamma_1 \left(\tau_1 + \frac{1}{\frac{\beta'}{\tau_2 \beta} + \frac{1}{\tau_2 \tau_3 \beta} + \frac{1}{\tau_3}} + \frac{1}{\frac{1}{\tau_2} + \frac{\beta}{\tau_3 \beta' + 1}} \right) \quad (14)$$

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

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

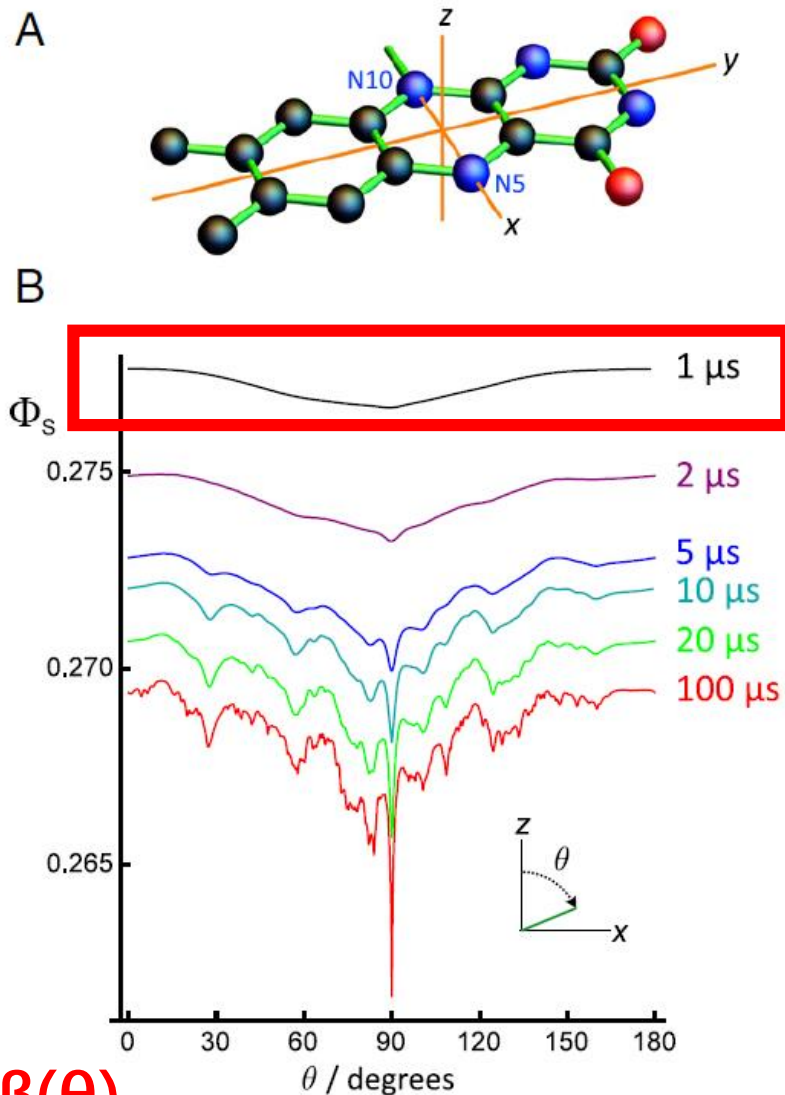
Simulation

Parameter initiation



- Singlet-triplet oscillation β 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_{photon}
 where $N_{\text{photon}}:C_1 = 10^{-3}$

Parameter initiation

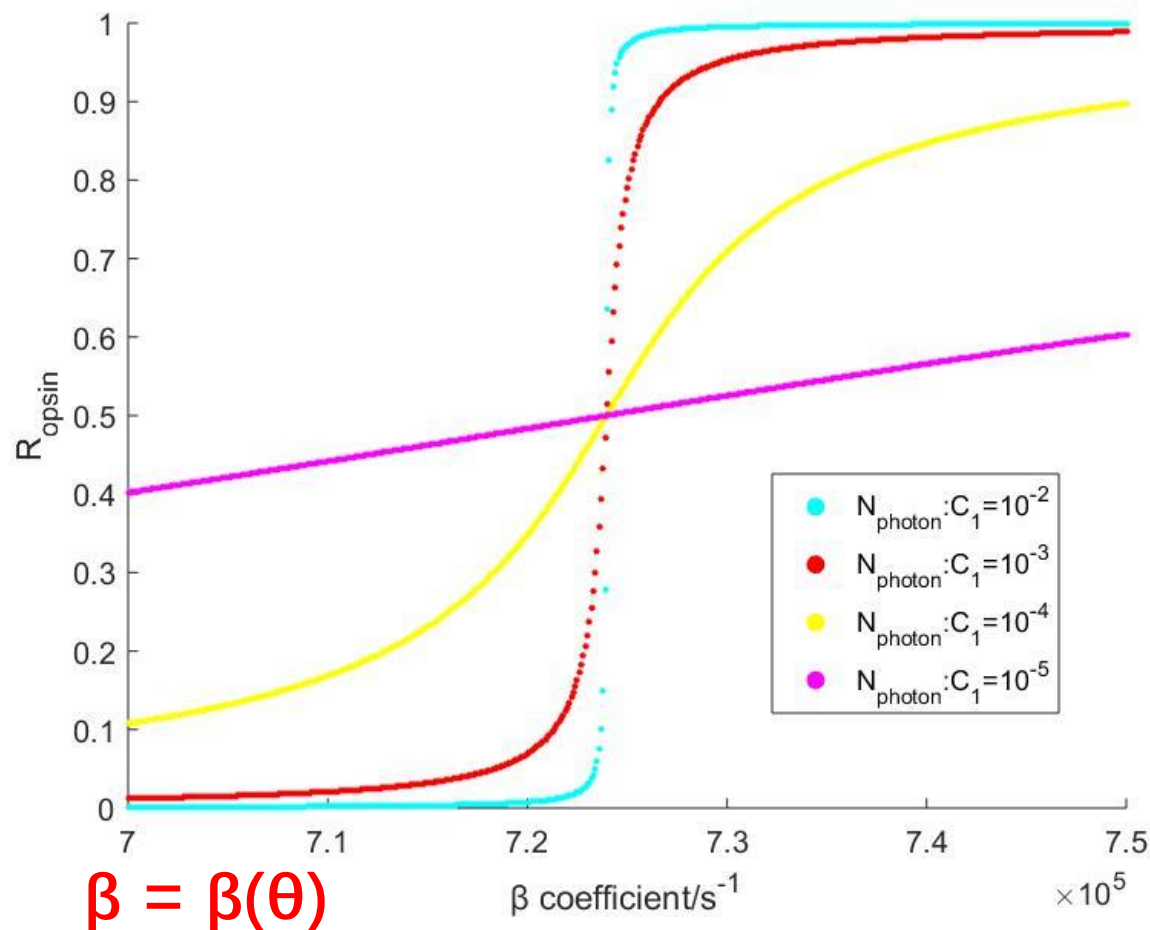


$$\beta = \beta(\theta)$$

- Singlet-triplet oscillation β 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_{photon}
 where $N_{\text{photon}}:C_1 = 10^{-3}$

Secondary amplification effect

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

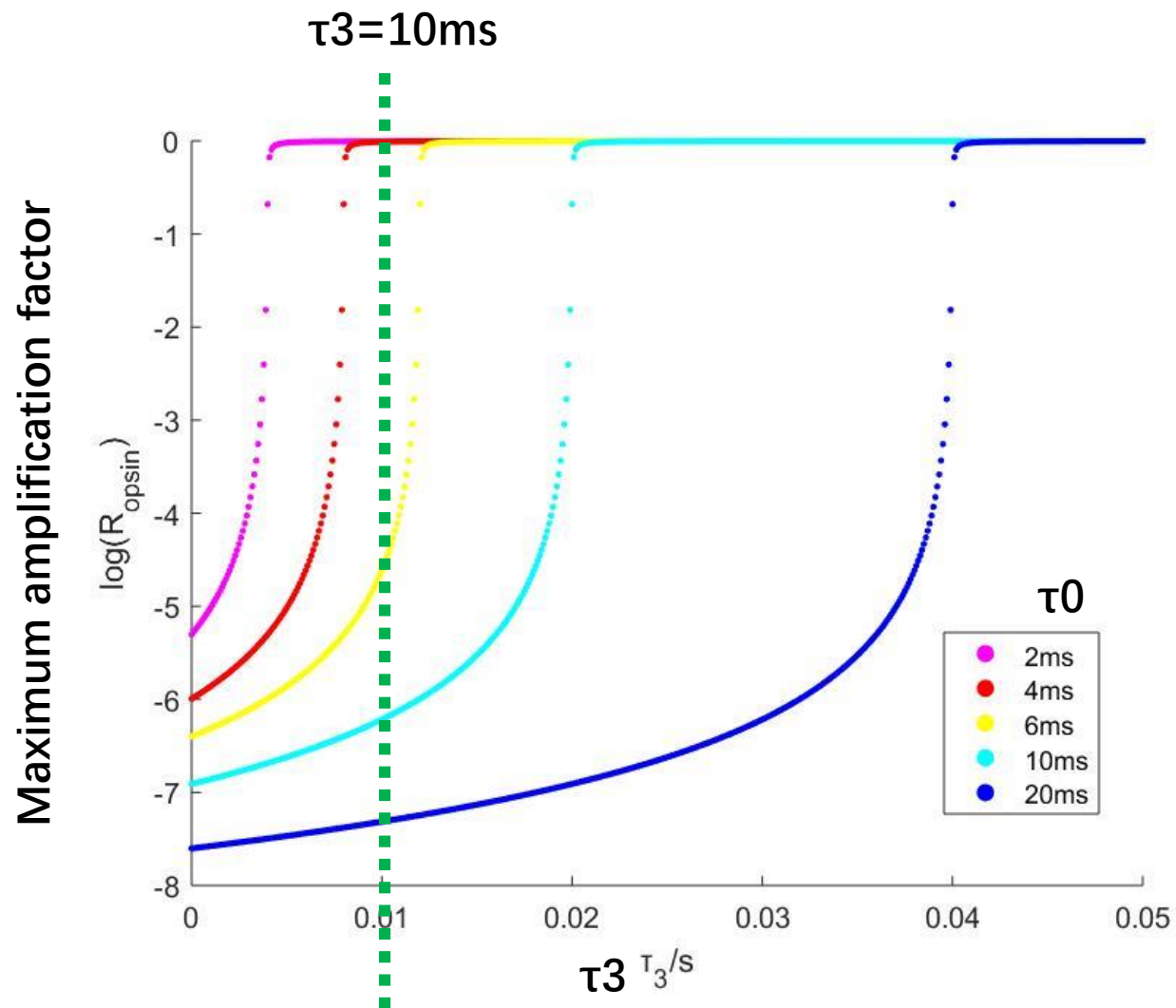


$$B = \tau_0\gamma_0 + 1$$

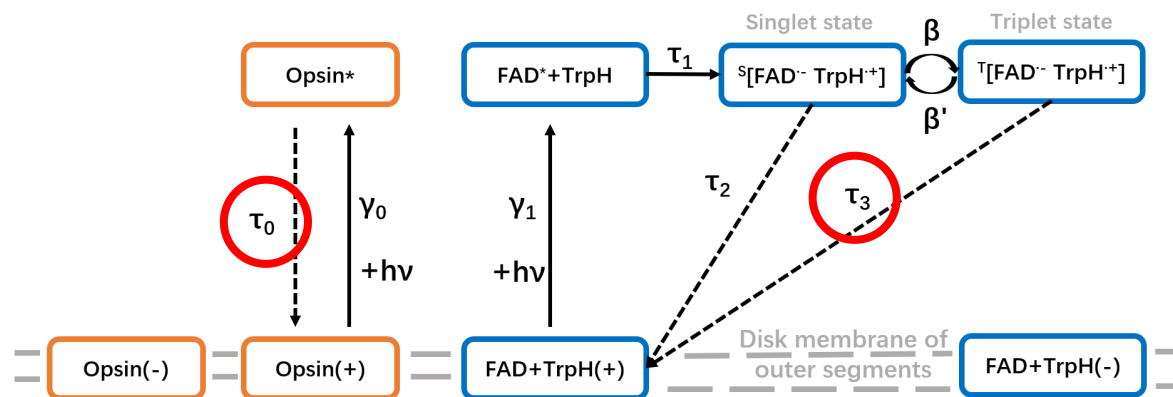
$$A = 1 + \gamma_1\left(\tau_1 + \frac{1}{\frac{\beta'}{\tau_2\beta} + \frac{1}{\tau_2\tau_3\beta} + \frac{1}{\tau_3}} + \frac{1}{\frac{1}{\tau_2} + \frac{\beta}{\tau_3\beta' + 1}}\right)$$

- $R_{opsin} \in (0, 1)$ representing the percentage of incident photons that are absorbed by rhodopsin.
- Ratio of incident photons and protein numbers are critical for amplification **indicating a upper threshold** for magnetoreception.

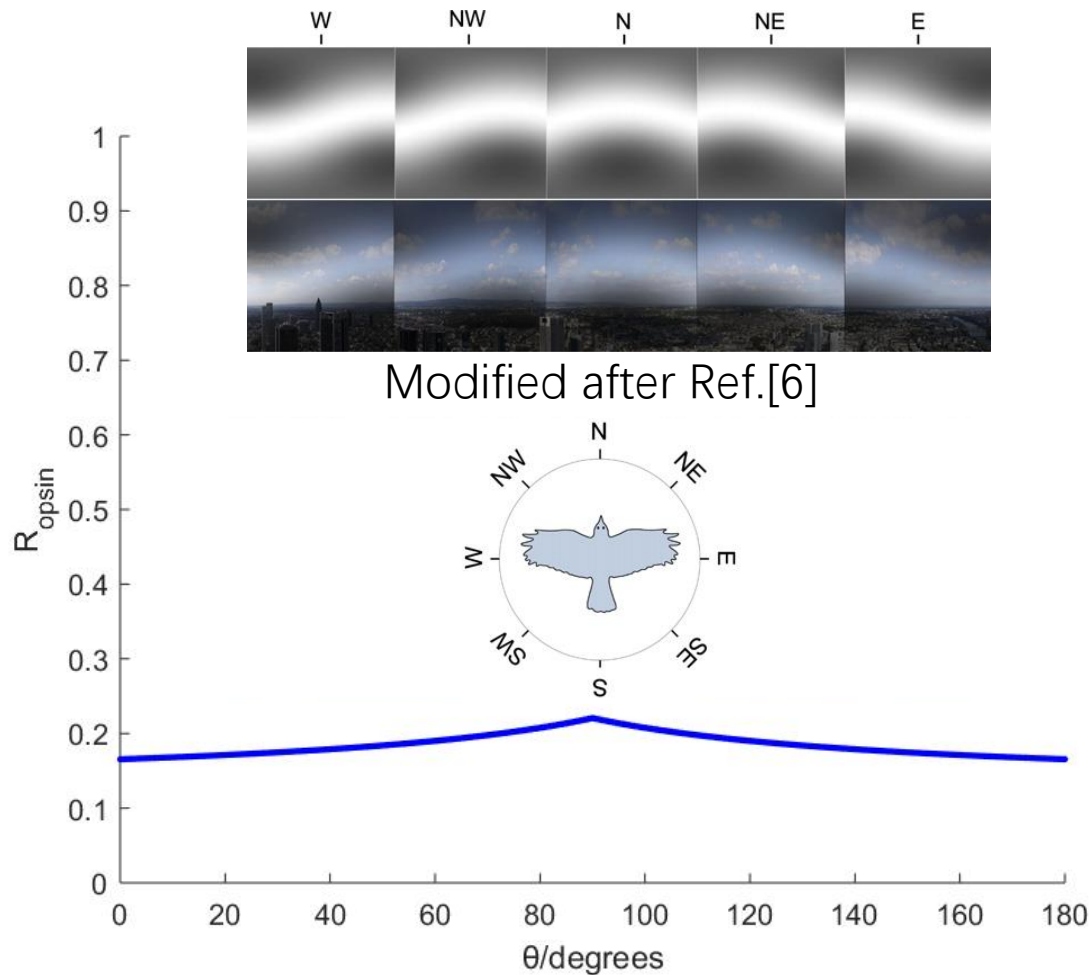
Secondary amplification effect



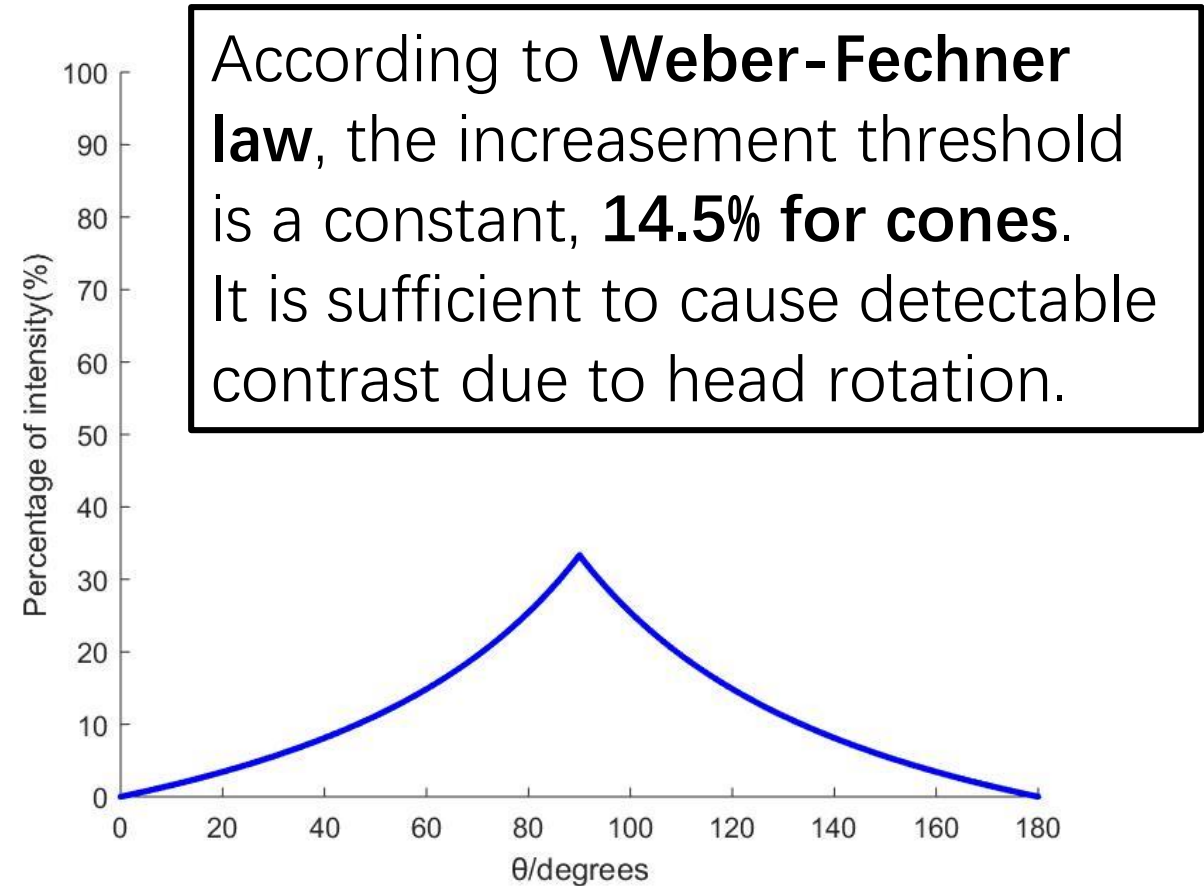
- The value of τ_0 is critical for the amplification effect.
- The secondary amplification effect is a key step to convert a physical cause into biological consequence.



Photon competition simulation result



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



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

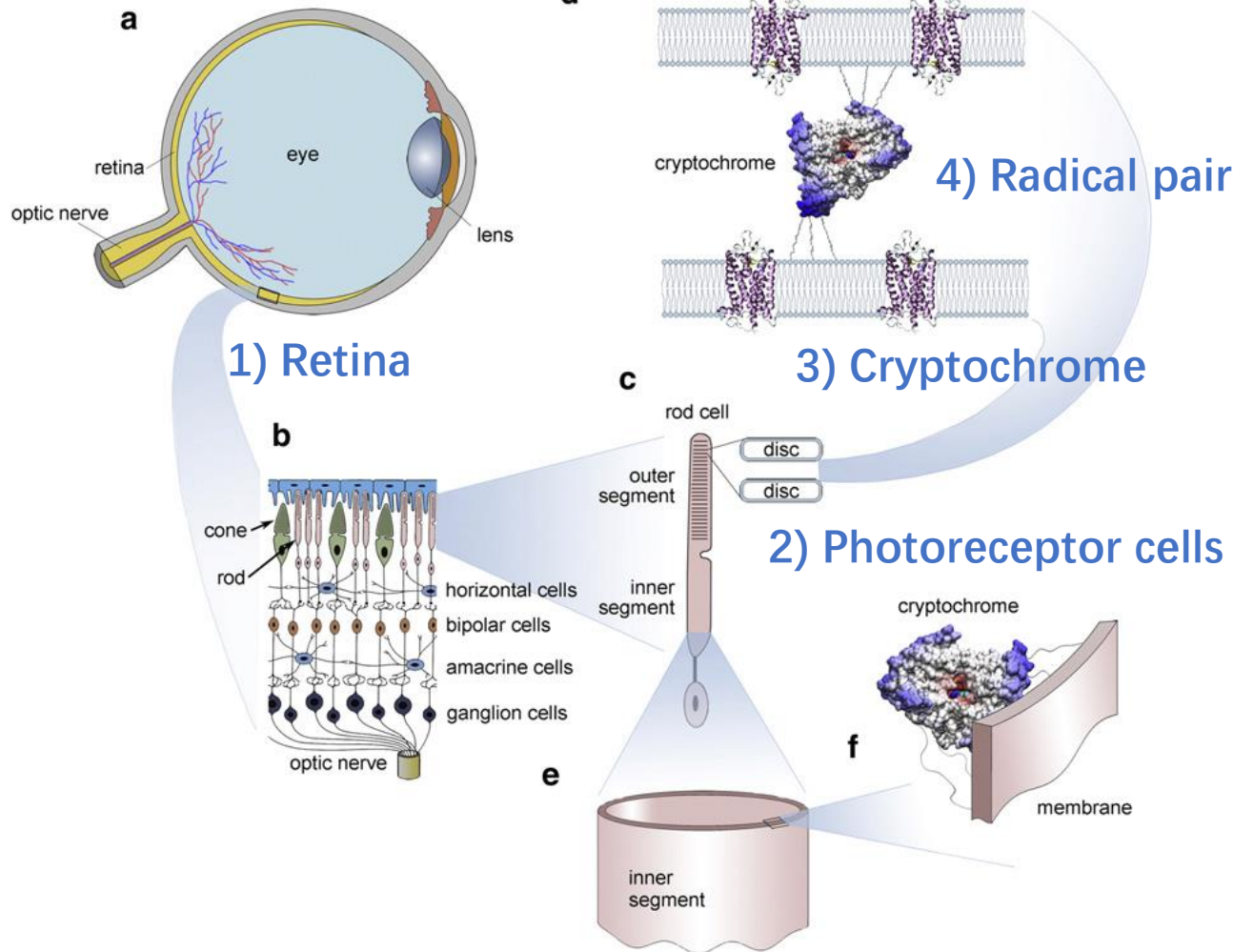
Experimental verifications

- Three strategies to verify OCC model:
- (1) Incident photon threshold for magnetic sensing.
- There is a upper photon threshold 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?

5) Visual pathway



- 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).

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 percept 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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Acknowledgement

- Thanks for Dr. Jiansheng Wu for the helpful instructions in theoretical realization of the biophysical model and warm guidance during the project.



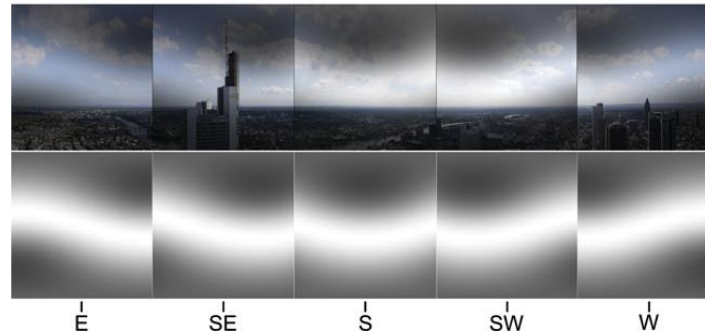
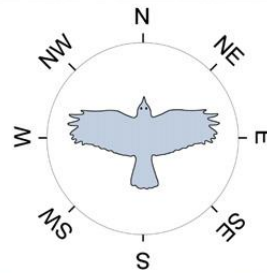
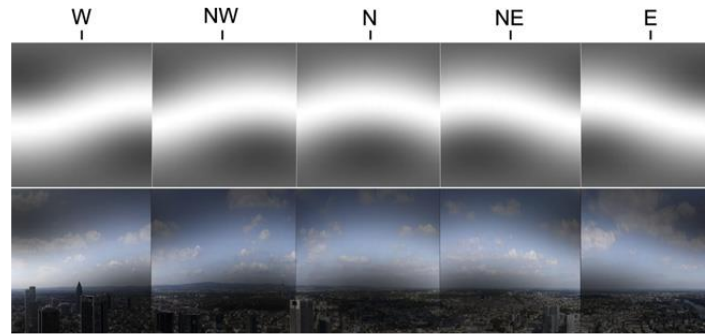
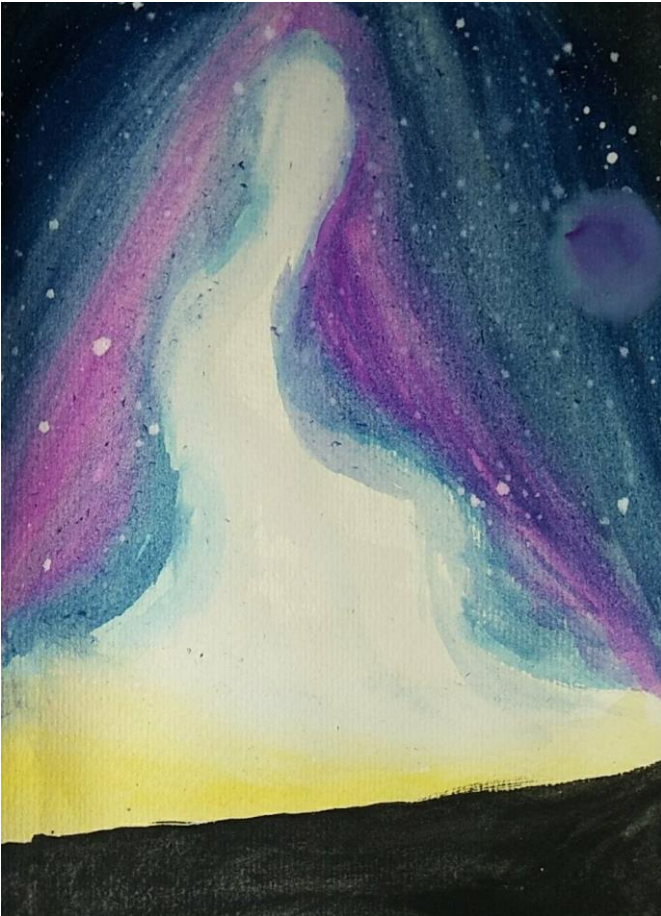
Jiansheng Wu,
Assistance professor
Department of Physics
SUSTech



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



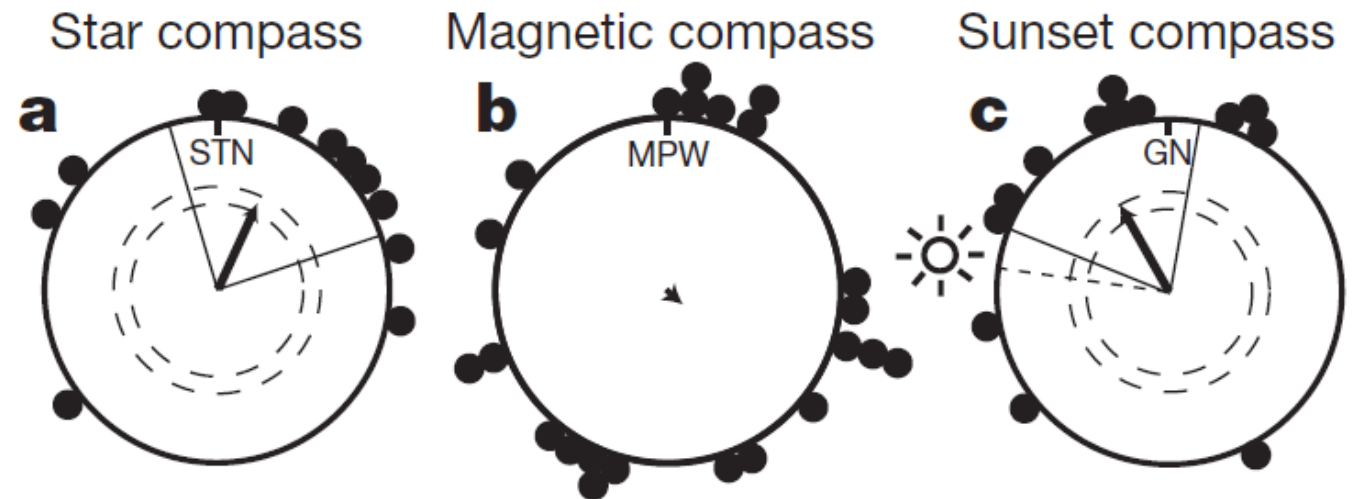
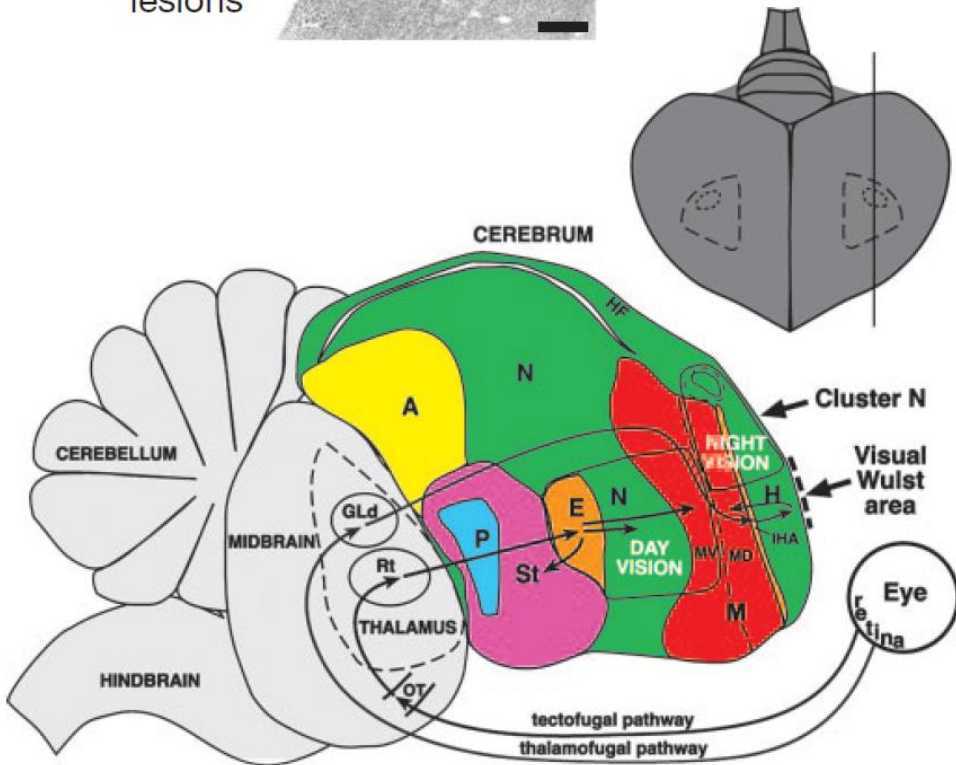
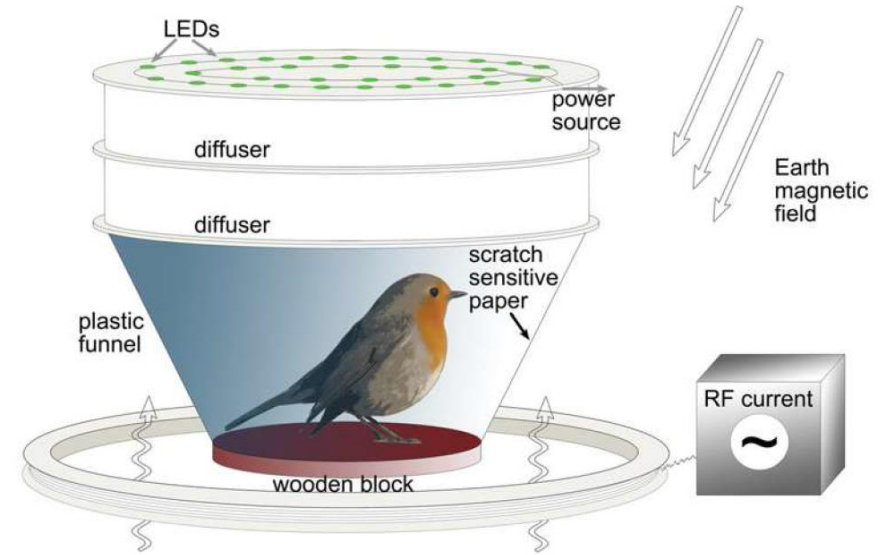
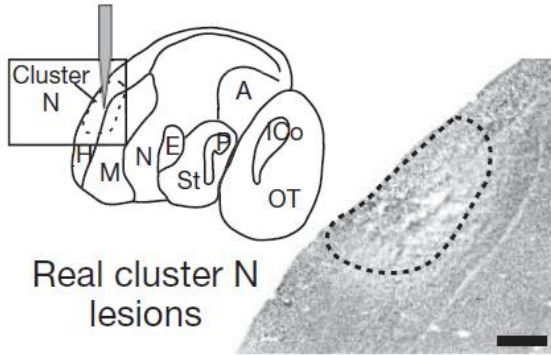
Thank you



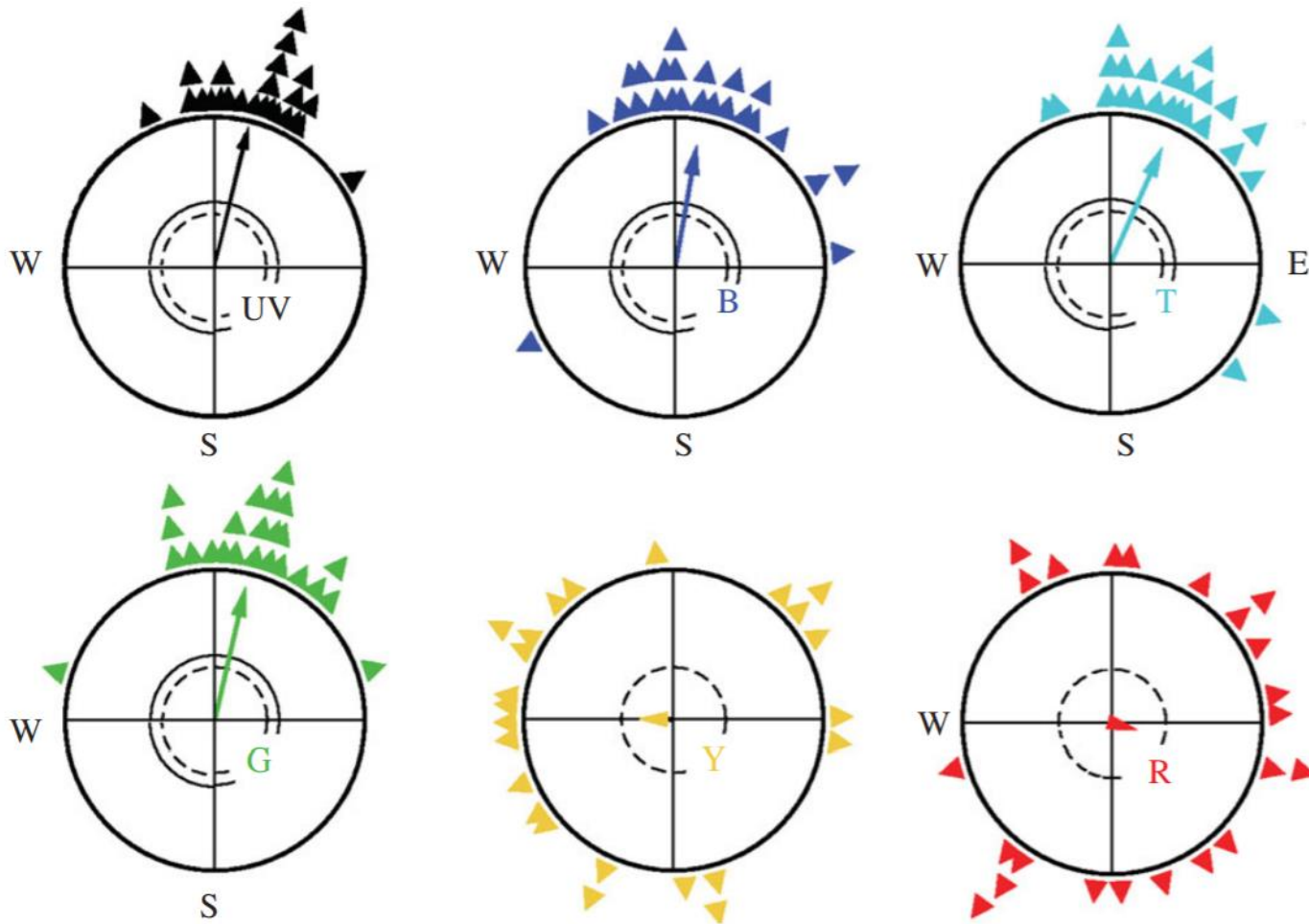
Appendix

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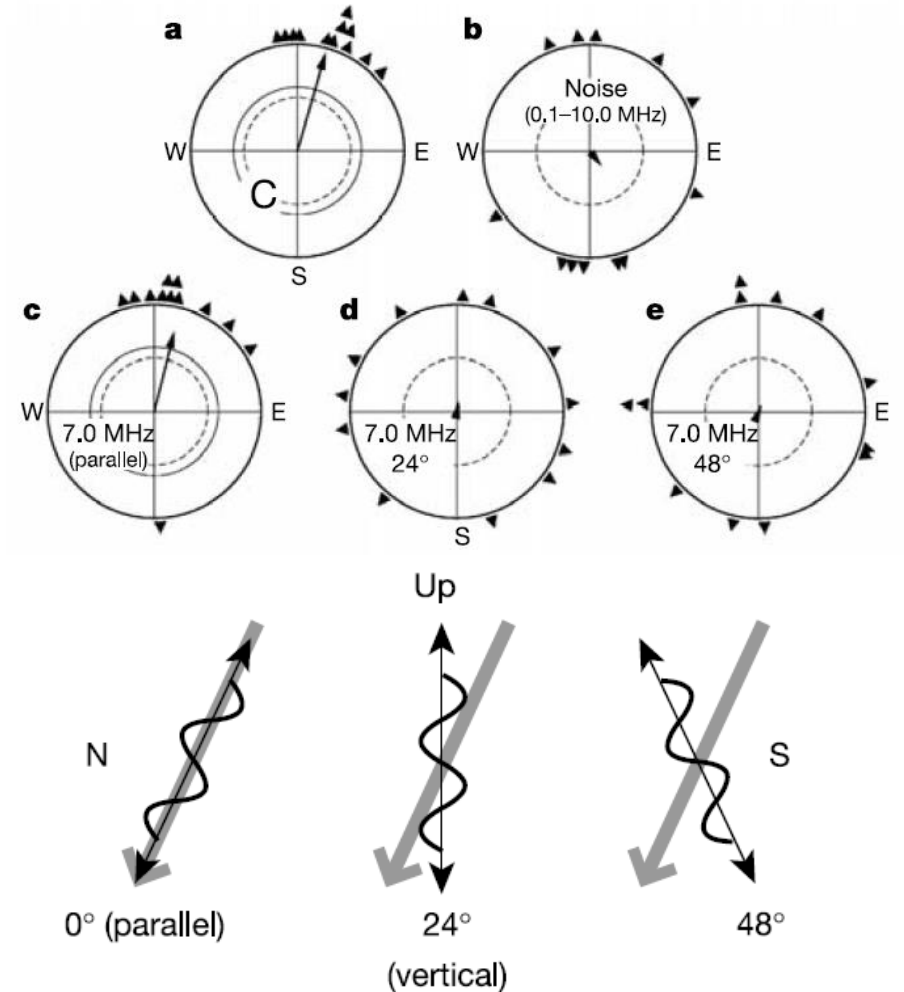
- 1) A neural connection between night vision and magnetic sensing in European robin.



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

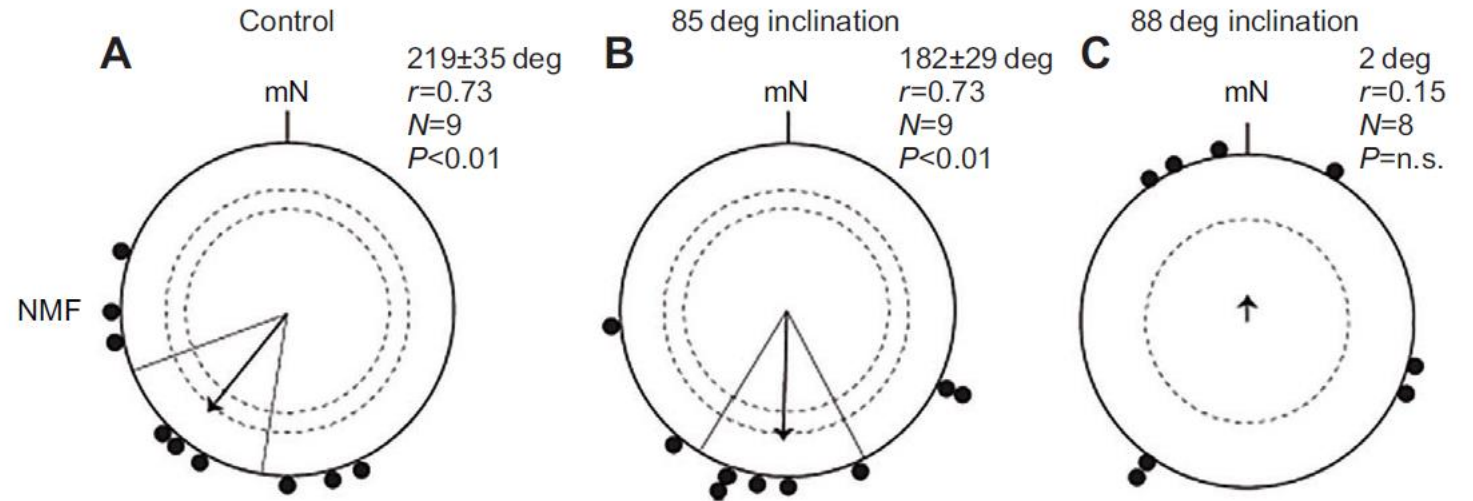
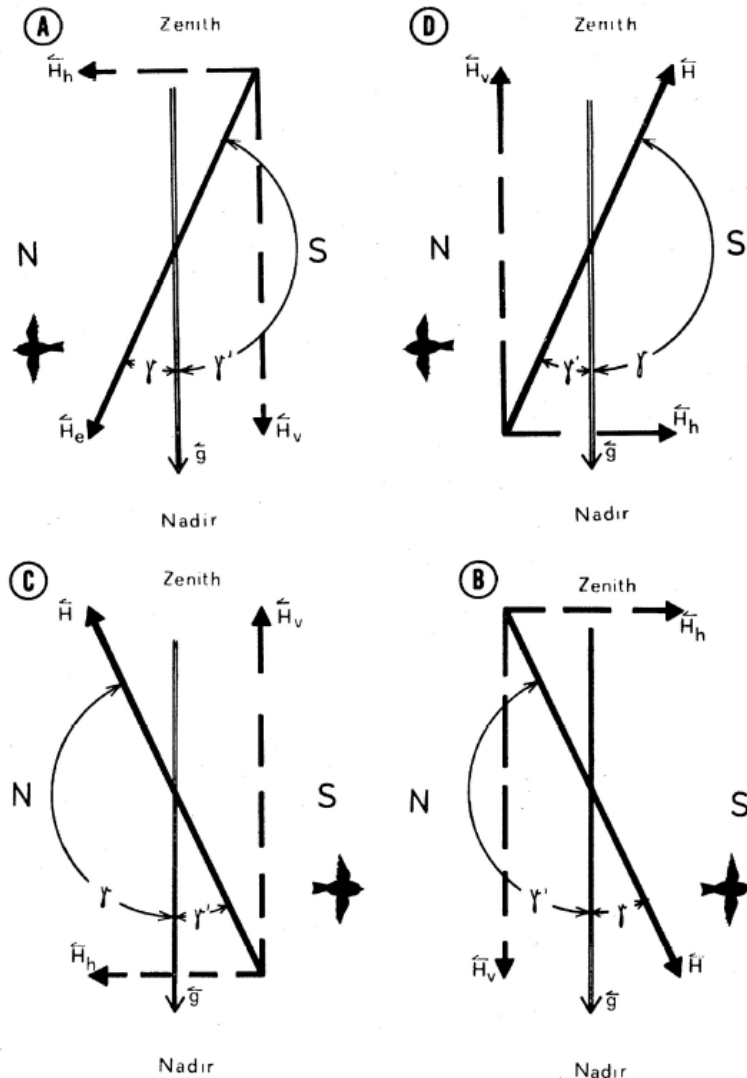


Orientation of European robin under monochromatic light of different wavelengths



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.

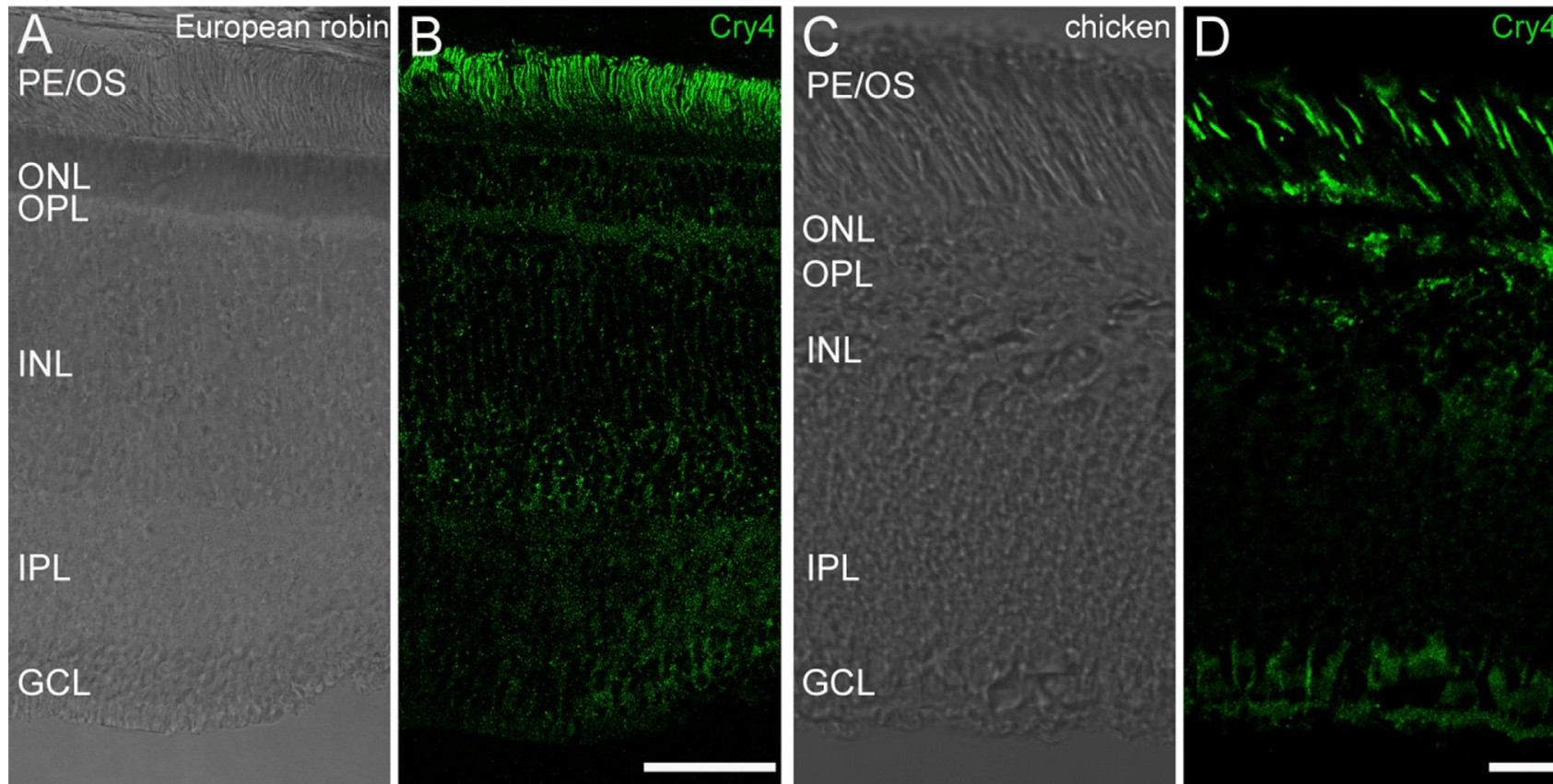


The orientation of individual blackcaps was tested in magnetic fields with 67, 85 or 88 degree inclination

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