O detetor infravermelho da Cascavel

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Café com Física São Carlos 2014

Colaboradores: Ifsc: J. Slaets, R. Pinto, L. Almeida, I. Zucoloto, R. Batista

Unesp Rio Claro: G. Gomes, D. Andrade; UFPe : P. Carelli.

Experimentos: Unesp Rio Claro

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Typical pit organs

Infrared Organs of Snakes: J. Herpetology, 45, No.1, 2011., R. C. Goris (2011)



Fig. 1. Typical snakes possessing pit organs. (A) Python molurus, albino (Pythonidae). (B) Corallus caninus (Boidae). (C) Gloydius blomhoffii (Crotalinae, terrestrial). (D) Trimersums stringeri (Crotalinae, arboreal). The arrows in C and D point to the pit. Modified from Goris et al., 2010. Channear block and the Targener (Crotalinae) arboreal). The arrows in C and D point to the pit. Modified from Goris et al., 2010.

Goris (2011) / /

Anatomy

Pit organ innervation



Fig. 3. Intervation of the crotaline pit organ. The pit membrane (red) is intervated by three bearches of the tingenital nerve (velow). Altherizations C, crebelum, E, evelual (bate); C, neticor oditions; MO, medial oblogate, N, notelli geneti, OS, oditatory table; OJ, opie etum, P, pit S, Sgandou CM, Vigenital gandiger, V, ophitaline transch et de the ingenital (Q, is pagnical). Zanadibar bench of the trigeminal (gr.) is gandios; Via, deep hanch of the maxillary transch; Via, supericial herench of the maxillary transch; Via, mandrabar bench of the triggminal (gr.) is gandios; Via, deep hanch of the maxillary transch; Via, supericial herench of the maxillary transch; Via, mandrabar bench of the triggminal (gr.) is gandios; Via, deep hanch of the maxillary transch; Via, and Katel the position of formation in the central bones (pot shown) where the rever turn test or sub rev should. Modified from Garsis et al., 2011: Color repositions of sported by the Thimas Burus; Final

Chegando ao Teto Optico



Figura 15 - Passos de cinurgia efetuada na cascavel. A) Com a cobra entubada retina-se a pele do topo da cabeça. 8) Fixa-se sua cabeça através de uma haste no coso entre os olhos (com resina de dentista). (C e) (Espocija do tete óptico, utilizando uma broca de dentisa, com a retinada de uma parte do soso do cráno.



Figura 16 - A) Teto óptico mostrando através da cabeça da cobra B) Teto óptico sendo mostrado através do microscópio; eletrodo penetrando nele.

Dissertação R. Batista

Pit organ cross section

Goris (2011)



2. Degrammic cross-sectional mappings of stake infrand record regars. Haldings indicates the location of the receptor terminals, (A) therebarg the is (g), constrolly. The receptors is not care to merginal and and dials of set (babilis care). (B) A bability the is (g), cantains, The receptors are location is rescaling and and dials of set (babilis care). (B) A bability the is (g), cantains, The receptors are location is relatively and the period relation of the system of the receptor set requestion is the period relation of the print system of the set relation of the print system of the print syst

Hartline 1982: Some 7000 thermosensitive

nerve endings of trigeminal sensory axons

on the 15 μ thick membrane with area 30 μ^2 .



Fig. 7. The pinhole camera effect. An infrared source throws the shadow of the pit mouth onto the membrane receptors, exciting the receptors one after another as it moves, creating a unique series of action potentials, which the central nervous system interprets as a pattern. Modified from Goris et al., 2010.

TNM = Terminal Nerve Masses, Goris(2011)

Estrutura superficial e direcional da membrana



Fic. 9. Illustration of how the surface architecture of the pit organ shown in Figures 8 makes the organ selective for intrared radiation. The shown in Figures 8 makes the organ selective for intrared radiation. The simulate the receptors (black dots) but reflecting away the shorter wavelengths of intrared radiation (red lines), allowing them to the micropits on the surface of the domes from the inner chamber back wall (Fig SC) and, further, by the micropits on the inner surface of the utility of the strategies of the domes from the inner strates of the but these will be absorbed in the light trap formed by the clustered large and small domes of Figure 8C and illustrated in the enlargement. Modified from Carie et al., 2010. Color reproduction supported by the





(Goris-2011)

-Membrane Properties

Electron micrograph of single TNM

Pit Capillary System - Goris (2011)

KED RECEPTORS



Fig. 4. A scanning electron micrograph of a single TNM viewed from the inner chamber. The pit membrane contains several thousands of these arrayed in a single layer just beneatin the outer epidermis of the membrane. Lower arrow indicates the single myelinated nerve fiber that terminates in the TNM. Upper arrow shows the point where the nerve loses its myelin sheath and branches out to form the nerve mass. Bar $= 10 \, \mathrm{um}$. From Goris et al., 2010.



Frc. 10. The capillary bed of a pit membrane (Gloydius blomhoffit), The blood vessels have been visualized by perfusion with India ink. Each capillary loop encloses a finite number of TNMs. In actuality the membrane is concave; slits have been cut around the edges to make it lie fat. R, rostral; D, dorsal. Bar = 250 um.

— Membrane Properties

Electron micrograph crossection of pit membrane (2010)



Fig. 13. A transmission electron micrograph of a cross-section through the pit membrane of a pit viper, Gluphias Neudoffii, showing how a capillar (selectisk) contacts two TMMs, presumably for coding them. The inset shows a magnified view of the dightly packed mitchendratis in a TMM. Abbreviations C, outer channels. First + JJM Modified from Gories et al., 2010.

- Membrane Properties

Table 1

Properties of biological infrared imaging and sensing organs and thermoreceptors

Animal/Insect	Threshold energy $(\mu W/cm^2)$	Threshold temperature change ($^{\circ}\!$	Distance to detection	Type of receptor	Directional sensitivity	Reference
Boid IR receptor (Boa constrictor)	176.89	0.003	16.4 cm	Specific warm	No	Barrett et al. (1970); de Cock Buning (1983); Molenaar (1992).
Boid pit organ (Python reticulatus)	59.75	0.026	28.3 cm	Non-specific warm bimodal (IR and tactile)	Yes	Barrett et al. (1970); de Cock Buring (1983); Molenaar (1992)
Crotaline pit organ (Agkistrodon rhodostoma)	10.75	0.003	66.3 cm	Specific warm	Yes	de Cock Buning (1983); Molenaar (1992)
Beetle antennal thermoreceptors (Melanophila acuminata)	May detect fires at short range	2.0	N/A	Warm receptor	Yes	Evans (1964)
Beetle pit organ (Melanophila acuminata)	60-500	0.01	60-100 miles	Warm receptor	Yes	Schmitz and Bleckmann (1998)
Common vampire bat pit organs (Desmodus rotundus)	50	-	8–12 cm	Warm receptor	Yes	Kurten and Schmidt. (1985); Molenaar (1992)
Blood-sucking insect antennae (Itratoma infestans)	-	Orient toward objects between 30 and 32°C	4-10 cm	No receptor identified yet	Yes	Lazzari and Nunez (1989)
Butterfly wing vein thermoreceptors (Troides r. plateni)	-	Respond to heating rate of 2.4-4°C/s	N/A	type II receptors (warm)	Yes	Schmitz and Wasserthal (1993)
Butterfly antennal thermoreceptors (Troides r. plateni)	-	May measure ambient temperatures	N/A	Type I receptor (warm)	Yes	Schmitz and Wasserthal (1993)

- Measurements

Recording from Optical Tectum



FIG. 5. Typical action potentials recorded from infrared neurons in nakes possessing infrared organs, in this case from the optic tectum of Gloydius brevicaudus. (A) Spontaneous discharge. Infrared neurons onstantly produce action potentials at irregular intervals, caused by andom stimulation by objects in the visual field of the pit all of which, by the laws of physics, emit infrared radiation of varying strength. (B) timulation by an infrared (830-nm) laser. The neuron responds with a burst of firing, with a varying degree of latency depending on the ndividual neuron. This is part of the encoding, which eventually produces a conscious image in the central nervous system. The solid ine shows the onset, duration, and cessation of the laser. A and B are rom the same neuron. (C) Response to a cold object (a popsicle). Arrows indicate the points at which the popsicle entered and left the pit's field of view. The spontaneous discharge at left ceases abruptly and responds with a strong burst when the stimulating object leaves he field of view. This shows that the infrared neuron can respond to an bject whose temperature is lower than the background radiation, for example, a wet frog. The scale refers to all records. Modified from Goris

(a)

- Measurements

Hartline 1983 - IR and Visual Space

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Directional Sensitivity - Bakken 2012





Fig. 8. Plot of directional sensitivity of the facial pit of C. *atrox*. Along both the horizontal (blue) and vertical (red) transacts the facial pit exhibits clear directional sensitivity. Threshold (2-axis) represents the greatest distance (up to a maximum of 30 cm) between emitter and facial pit that generated a significant neural response of background activity.

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- Measurements

Water flowing on Trigeminal Nerve - Bullock 1956, $\Delta T \sim 0.025^{\circ}C$



Text-fig. 4. Sensitivity measured by change in temperature. Water flowing at constant rate over the pit membrane is electrically heated from a certain moment. Three different rates of temperature rise, estimated by reading thermcourple records at 0.3, 10 and 2.0 sec. Response measured as interval between successive spikes except at high frequency, when the average interval for two or three successive spikes is pitcled. The upper, middle and lower hand drawn curves (circles, crosses and dots, respectively) correspond to the stimulus curves in the same relative positions. There is uncertainty in the position of zero time on the record, owing to the small thermcourple defixion, of about 0.1 sec. Nevertheless, the response is very abrupt and permits estimation of the ΔT approaching threshold. Ordinate at right in ^o C.

Bakken & Krochmal(2007). Thus, the membrane may conceivably respond to contrasts of less than 0.001°C (Recording response threshold from Trigeminal Nerve). The mechanism by which such sensitivity might be

obtained is presently unknown

- Measurements

Pit

- innervated by two ganglia of the trigeminal nerve.
 Warm myelinated A-delta fibers from the Ophthalmic ganglion: innervate dorsal part of membrane, Maxillary ganglion: two bundles innervate the ventral and rostral areas of the membrane.
 Terminals of nerves from these ganglia innervate unevenly the pit membrane (~ 600 fibres),but each has its own *territory* within the membrane.
- 2 Pit membrane is also innervated by scores of unmyelinated C fibers, some of which are probably nociceptive fibers, whereas others may belong to the autonomic (sympathetic and parasympathetic) nerve system controlling the blood vessels of the membrane.
- 3 Sensory receptors of all kinds contain a more-or-less large number of mitochondria, but the mitochondria in the snake TNMs surpass by far the numbers in other known sensory organs. It may be that these massed mitochondria form part of the mechanism, whereby the pit receptors respond extremely rapidly to minute temperature changes, but this has yet to be proven.

(Goris 2011, Goris and Nomoto, 1967; Goris and Terashima, 1973; Kishida et al., 1980; Berson and Hartline, 1988).

-Membrane Properties

Molecular structure

<u>Gracheva et al.</u>: Molecular basis of the infrared detection by snakes. Nature 464:1006-1012, 2010. Pit membrane:

Passive antenna for radiant heat,

transducing thermal energy to heat-sensitive channels on embedded nerve fibres.

 \rightarrow Snake TRPA1 is a heat-activated channel.

TRPA1 roperties:

- inactive at room temperature,
- but robustly activated above $27.6 \pm 0.9^{\circ}$ C.
- no response to cold (12°C).



Figure 3 | Functional analysis of snake TRPA1 channels. a, HEX:93 cells expressing cloned ratticsnake or rat snake TRPA1 channels were analysed for heat or mustared of 200 pM AITC2 eX⁻¹ Co-vocket responses using calcium imaging: colour bar indicates relative change in fluorescence ratio, with purple and white denoting the lowest and highest cytoplasmic calcium, respectively (n= 20 cells per channel). B, Relative heat response profiles of

rattlenake and rat snake channels expressed in oncytes (response at each temperature was normalized to the maximal response at 45° C; holding potential (V₁₁) = -80 mV; $n \ge 6$). Duta show mean " $\pm d$. C, Arbenius plots show thermal thresholds and Q₀ values for baseline and evoked responses of rattlensnke (Left) and rat snake (right) TRPA1 channels, as indicated (temperature ramp of 1°Cs⁻¹).

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-Membrane Properties

Firing Patterns

The firing patterns of neurons associated with pits at all levels of the nervous system (peripheral nerves, medulla oblongata, optic tectum, nucleus rotundus of the thalamus, anterior dorsal ventricular ridge of the telencephalon; for details, see below) have three common characteristics.

(Goris 2011, Goris and Nomoto, 1967; Goris and Terashima, 1973; Kishida et al., 1980; Berson and Hartline, 1988).

- The neurons of other sensory modes (e.g., lateral eye vision) are silent. In contrast, infrared neurons fire constantly at irregular intervals. All objects, constantly give off infrared radiation generating *spontaneous discharge*. At a body temperature of 25° C, frequency is 10-30 spikes/sec (varies with species, body temperature, and individual neurons).
- 2 Firing increases in response to any stimulus of higher temperature (or stronger IR radiation) than the background. Firing frequency changes with IR λ of the stimulus, strongest at

 $\lambda = 3 - 15 \mu m$

- = wavelength normally radiated from the body surface of endothermic animals
- **3** For temperature lower than that of the background \rightarrow firing frequency decreases (e.g., Bullock and Diecke, 1956; Goris and Nomoto, 1967; Fig. 5C).

Black Body Radiation,Flux, Geometry, Background and Target

Um pouco de Física

Radiação do Corpo Negro.

Radiation by unit volume and unit solid angle				
$1\ cal = 4.1868\ Wsec$	$k = 1.381 \cdot 10^{-23} \ J/^{\circ} K$			
$c=3\cdot 10^8\ m/sec$	$\hbar = 6.600 \cdot 10^{-34} \ J \cdot s$			
$T_0 = 273.16^{\circ} K$	$\sigma = 5.6522 \cdot 10^{-8} \frac{W}{m^2 \cdot K^4} .$			
Spectral densitiy $[E/V/d\Omega]$ is				
	$e_{0}(\omega) = \frac{1}{V} \frac{2 \cdot 4\pi}{(2\pi)^{3}} \frac{dE_{\omega}}{d\omega} = \frac{\hbar \omega^{3}}{4\pi^{3} e^{3} (e^{\hbar \omega/kT} - 1)}$			

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Black Body Radiation, Flux, Geometry, Background and Target

Radiation by hemisphere

Emitted flux by surface \hat{n} : $J(\omega, \theta) d\Omega = ce_0(\omega)d\Omega = ce_0(\omega)$ Emitted flux falling on unit surface S tilted by $\hat{n} \cdot \hat{n}_S = \cos \theta$: $J(\omega, \theta) \cos \theta d\Omega$

Total emitted flux falling on S:

 $J_0 = c \int_0^\infty e_0(\omega) d\omega$

Integration over frequencies yields

$$\int_{0}^{\infty} e_{0}(\omega) d\omega = \int_{0}^{\infty} \frac{\hbar \omega^{3} d\omega}{4\pi^{3} c^{2} (e^{\hbar \omega / kT} - 1)} = \frac{\hbar}{4\pi^{3} c^{2}} (\frac{kT}{\hbar})^{4} \int_{0}^{\infty} \frac{x^{3} dx}{e^{x} - 1} = \frac{(kT)^{4}}{4\pi^{3} c^{2} \hbar^{3}} \frac{\pi^{4}}{15}$$

integration:

$$J_0 = rac{(kT)^4}{4\pi^3 c^2 \hbar^3} rac{\pi^4}{15} = \sigma T^4$$
 (Stefan-Boltzmann)

with $\sigma = \frac{\pi^2 k^4}{60\hbar^3 c^2}$.

For conical radiative aperture with angle θ , instead of emitting half-sphere: $J_{\Omega}(\theta) = \sigma T^4 \sin^2 \theta$.

Energy Flux: experimental results

Energy Flux: experimental results

Previous Experimental Results from nerve recordings

1 Bullock(1956);

> Trigeminal nerve recording and supposing all IR radiation is absorbed: Threshold flux $\Delta \Phi = 75 \ \mu W/cm^2$;

Single fiber recording with water flowing on pit membrane:

 $\Delta T_{min} \sim 0.003^{\circ}C$

These figures cannot be given great reliability because this sensitivity pushes to the limit the usefulness of the available temperature recording system and we are not sure of the form of the initial change in temperature.

2 Goris et al.(1967): IR-laser $\Delta \Phi \sim 100 \ \mu W/cm^2$ at membrane.

Experimental Results using $\simeq \sigma (T_1^4 - T_2^4) (R/D)^2$

1 Buning et al.(1981): Shutter-Signal independent of background, record from midbrain tectum $\Delta \Phi = 10.76 \ \mu W / cm^2$.

```
2
    Ebert & Westhoff (2006):
     Shutter, Behavioral threshold for Crotalus atrox:
     \Delta \Phi = 3.35 \ \mu W/cm^2
     for D = 100 \ cm.
```

Energy Flux: experimental results



Figura 24 - Experimento sendo realizado com o braço mecânico e placa peltier.



Figura 22 - Foto do estimulo. Resistência circular acoplada no meio de arco de alumínio, dentro da cúpula de cobre (Todos devidamente pintados com tinta fosca preta).

Rio Claro, Dissertação R. Batista

Energy Flux: experimental results

Rio Claro results for ΔT

For a response threshold of $\sim 3^{\circ}C$, we get All wavelength:

 $\Delta T_{Min} \sim 0.01^{\circ} C.$

Assuming absorption only in the $\lambda = 4 - 15 \mu m$: $\Delta T_{min} \sim 0.005^{\circ} C!$



Energy Flux: experimental results

Numbers for our Experimental Setup

Total radiative heat flux is \sim time-independent!

For

IR background temperature $T_f = \mathbf{10}^{\circ} C$,

Moving target radiating area $A_a = 2 * 2 \ cm^2$,

Moving target temparture $T_a = \mathbf{13}^\circ C$,

Target to Pit distance $D = 10 \ cm_{\odot}$

 $f_{\lambda} \sim 1/2$

For IR-wavelenghts the irradiance contrasts are:

$$\delta \Phi = \Phi_{f+a} - \Phi_f = \frac{\sigma}{4\pi D^2} (T_a^4 - T_f^4) A_a f_\lambda = 9 \frac{\mu W}{cm^2}.$$

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Signal to Pit-background ratio:

- Signal Flux: $\delta \Phi = 9 \frac{\mu W}{cm^2}$.
- Heat flux from snake's pit: $\Phi_{pit} = \sigma T_{pit}^4 \epsilon_{pit} f_{\lambda} = 6650 \ \frac{\mu W}{cm^2}$

 $T_{pit} = 22^{\circ}C_{\gamma}\epsilon_{pit} = 0.3$

Include membrane absoprtion factor: $e_{abs} \sim 0.5
ightarrow 0.5$

$S = (\epsilon_{abs} \Phi_{pit}) / \delta \Phi \sim 350$

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Signal to noise ratio

Modelling a Historic Oil-Tank Fire Allows an Estimation of the Sensitivity of the Infrared Receptors in Pyrophilous Melanophila Beetles, Helmut Schmitz, Herbert Bousack (2012)

Beetles coming from 60 - 130 km



Figure 2. Identification of section 36 and the place of the old oilfields using [24,98]. doi:10.1371/journal.pone.0037627.g002

IR detector from *first* principles.

Problem: detect infra-red radiation embedded in infra-red background.

Nobody tells how heat accumulated in membrane is carried off! In the vicinity (~ $200\mu m$) of laser stimulation ($\lambda = 0.8\mu m$) get 50 % increase in blood flow in

 ~ 4.5 msec! (Goris 2007)

Equivalent problem in Vision:

Detect a light source embedded in an environment at temperature

 $T_{equivalent} \equiv \frac{\lambda_{infrared}}{\lambda_{visible}} = 10000^{\circ} K$

The whole Pit Organ would blaze like the sun! Is mechanical deformation of the pit an effective stimulus?

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Heating time/temperature dependence

Signal Flux $\delta \Phi = 9 \frac{\mu W}{cm^2}$.

Energy increase due to signal on membrane $\delta Q_{mbr} = \delta \Phi A_{mbr} = 1.15 \ \mu W.$

 $\begin{array}{l} \textit{Membrane absorption factor } \epsilon_{mbr} = 0.6 \\ \delta Q_{mbr} = (\epsilon_{mbr}c_vV_{mbr}\rho_{mbr})\delta T_{mbr} \text{ - data as for } \mathsf{H}_2\mathsf{O} \end{array}$

 $\begin{array}{l} \mbox{Temperature change:}\\ \delta T_{mbr} = 0.002 \ \mbox{C}^\circ/\mbox{sec} \end{array} \\ \mbox{Is there enough machinery to carry off the heat} ??? \\ Q_{pit}/\delta Q_{mbr} = 1090 \\ \mbox{What is the response-time of the membrane} \\ \sim 4 \ \mbox{msec (Goris 2007)!} \end{array}$

Where do we stand?

To do:

1 Study dependence on target movement & contrast sharpness.

- 2 Include geometrical factors and positional dependence of ϵ_{abs} .
- a Provide an ideal IR-detector in the presence of background and noise.

In contrast to statements by Bullock/Diecke (1956) and T. de Cock Buning (1983) arguing the

independence on snake temparature!

Show that the detctor is compatible with pit anatomy & physiology.

What is the thermo-transduction mechanism?

Future

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Effect of Snake Temperture - does it really cancel out?

Buning (1983) Considers background irradiation by snake's body Did experiment at ambient temperatures: $21C^{\circ}$ and $15C^{\circ} \rightarrow$ same threshold for response.

with and without chopper:

 \rightarrow snake-temperature cancels out in the difference.

(Assumes instantaneous reaction to changes in temparture gradient. Also: noise adds up!)

O que pode ajudar?

- Surface architecture
- Discovery of a Novel Accessory Structure of the Pitviper Infrared Receptor Organ (Plos1 2013), Carlos Jared et al., Departamento de Zoologia, Instituto de Biociencias, USP

Include heat lost/absorbed from membrane to air by convection

Amount of heat lost by conduction through the air to the pit walls:

Assume the snake being able to cool the membrane by

$$\delta T_{pit-mbr} = -0.1^{\circ}\mathrm{C}$$

Thermal conductivity of air: $k_{air} = \rho v c_p l_{mfp}/3 = 0.026 W m^{-1}/^{\circ} C$, Typical distances ~ 1 mm $Q_{Air} = [k_{air}(1/d_1 + 1/d_2)] * \delta T_{pit-mbr} A_{mbr} = 65.345127 \ \mu W$! Bakken & Krochmal 2007

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Shutter Experiment - Snake Temperature cancels

Originally this formula was designed to describe the net flux of radiation between two heat radiating objects with temperatures T1 and T2. Therefore, one tends to substitute for T1 the temperature of the pit organ (or the snake) and for T2 the temperature of the warm test object (or mouse/ral). However, in terms of physics the test situation described in this paper implies a change of radiant flux between before and after opening the shutter. Consequently, in the final formula the temperature of the snake is cancelled.

radiation density (cal/cm² · s) = $\frac{\sigma \times A}{\pi \times D^2}$ ((T⁴ stimulus - T⁴ snake) - (T⁴ shutter - T⁴ snake)) = $\frac{\sigma \times A}{\pi \times D^2}$ (T⁴ stimulus - T⁴ shutter).

In other words, the change of radiant heat influx on the pit organs can be regarded as the result of a change in the infrared environment, i.e., that small area which only changes in temperature (in this case the shutter and heat exchanger or the presence or absence of a mouse).

This implicates that the temperature of the sensor is not relevant for the ability of the system to discriminate temperature differences in the environment. Although, the intensity of the receptor response towards a specific stimulus is, of course, a function of a biologically related optimum temperature of the body (de Cock Buning 1981b, c: Hensel 1975), on the other hand, there is no argument from the physics to expect a relation between the threshold value and the temperature of the body. The same conclusion was reached by Bullock and Diecke (1956) and illustrated by one of their experiments. 'Experimentally, in a certain whole nerve preparation the response to a hand at 20 cm in a room equilibrated to 21 °C was about threshold. Wheeling the preparation and equipment into a cold room at 15 °C nearly doubled this distance. to 37 cm. There was no noticeable difference in threshold between the first moments when the snake's body temperature was still warm and spon-

Protein dynamics of vibrational states



Yamato(1998)

FIGURE 11.1 Mode diffusivity (black) and vibrational mode density (gray) computed for myoglobin. At low frequency, to about 100–150 cm⁻¹, where the vibrational modes are delocalized, the mode diffusivity is relatively large and becomes smaller with hiereasing frequency, in contrast to the vibrational mode density. Trends in the two quantities parallel one another at hietper frequency, where the vibrational modes are localized.

Photoreaction: in general, however, the functionally important motion of a protein along the reaction coordinate is largely masked by the thermal fluctuatios, making it difficult to analyze the effect of an external stimulus on the protein dynamics (*Heat Transport in Proteins*, Leitner 2010).

Future

É isso ai pessoal!

Obrigado!

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