

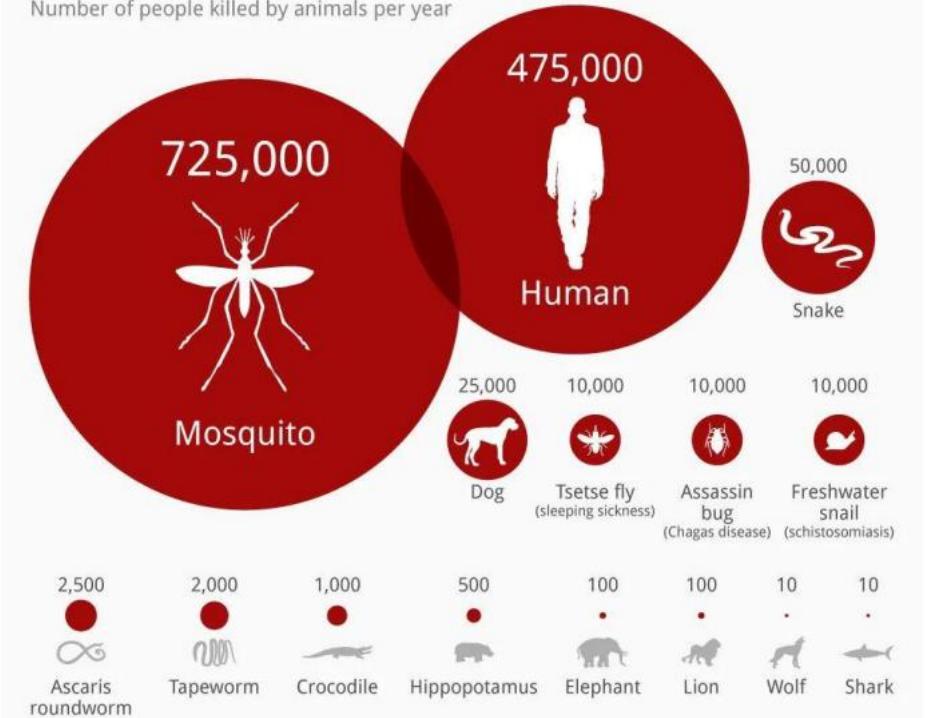
Don't bite me, mosquitoes!

李子奇、金思慧、姜思梅

2024.09.26

The World's Deadliest Animals

Number of people killed by animals per year



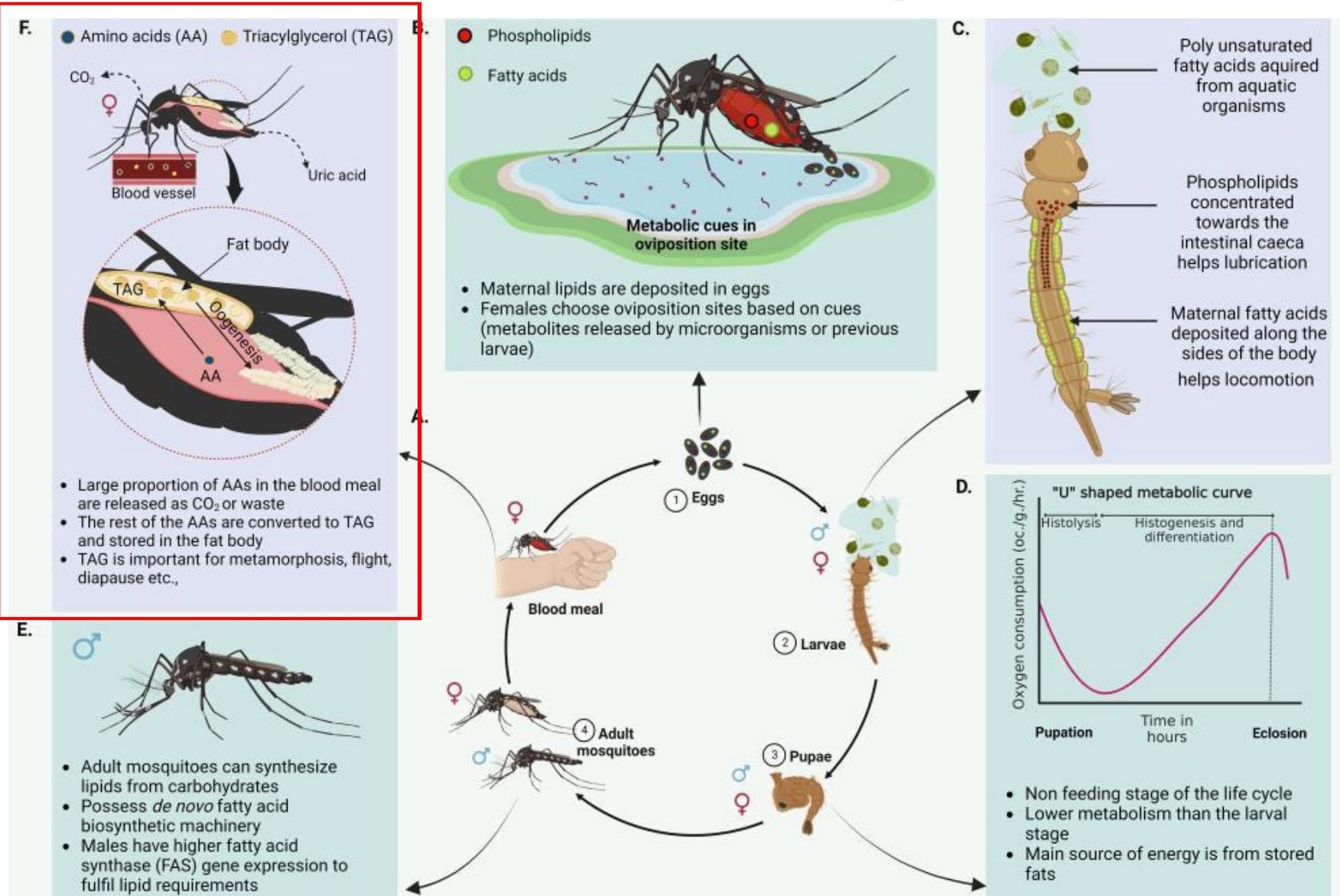
Content

- Overview of mosquito feeding——李子奇
- The internal mechanisms govern feeding——金思慧
- The external environment affects feeding——姜思梅

Overview of mosquito feeding

李子奇

The life history of the mosquitoes

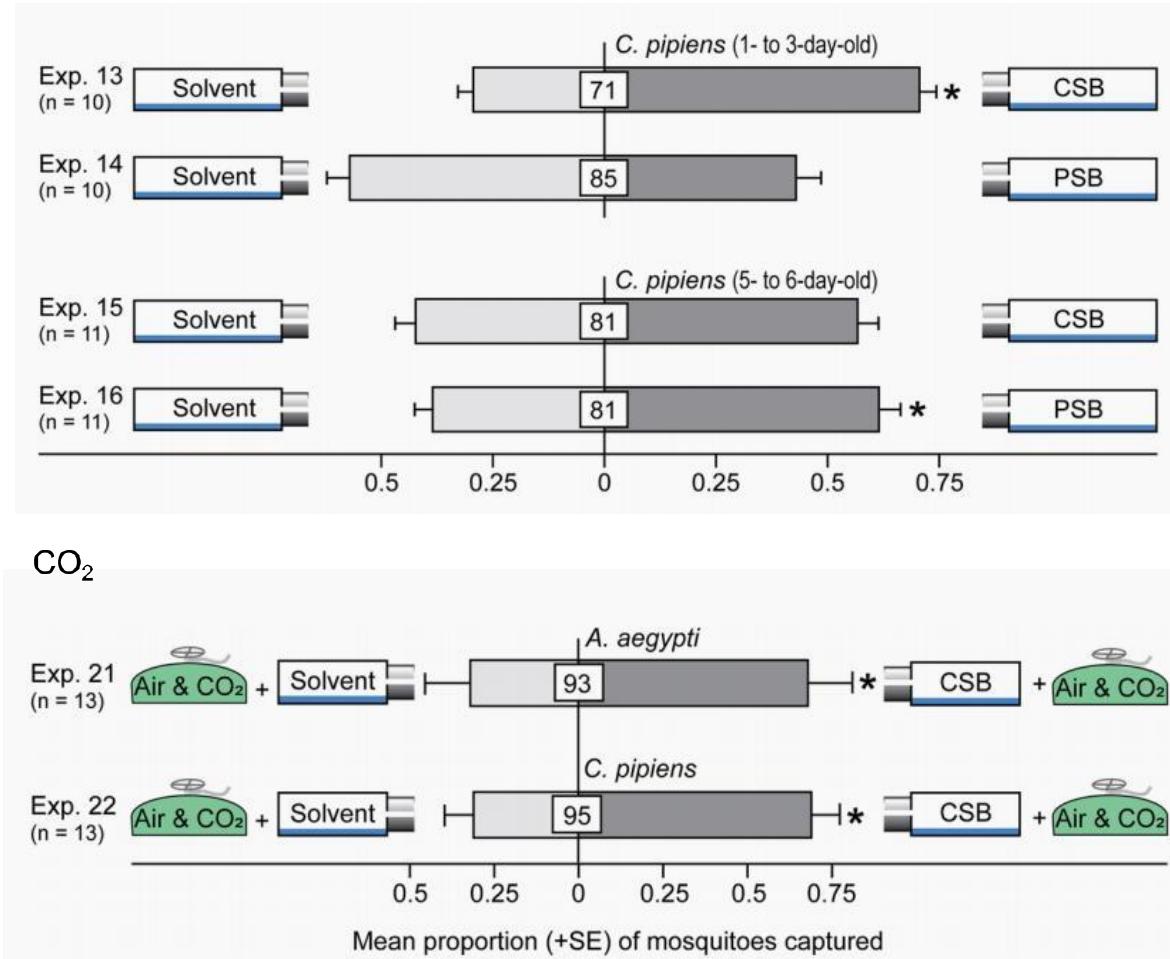
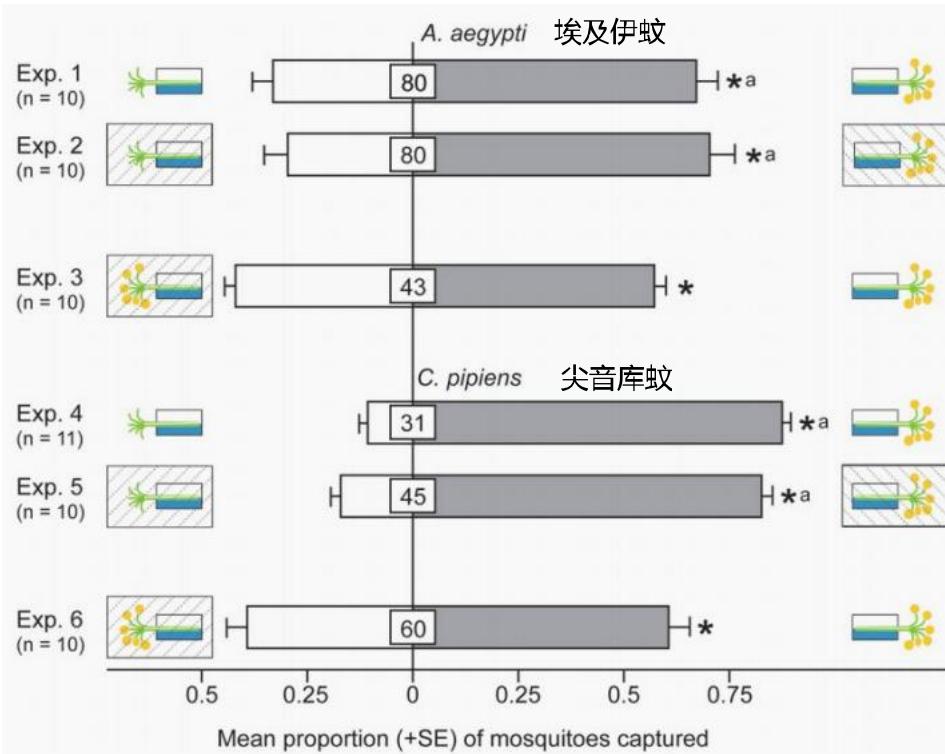
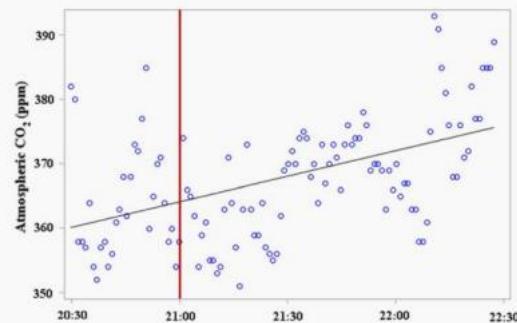


Ratnayake
OC,Chotiwana N,et al.,
Front Cell Infect Microbiol,2023

How do mosquitoes find nectar ?

- Visual cues
- Olfactory cues (CO_2)
- Gustatory cues

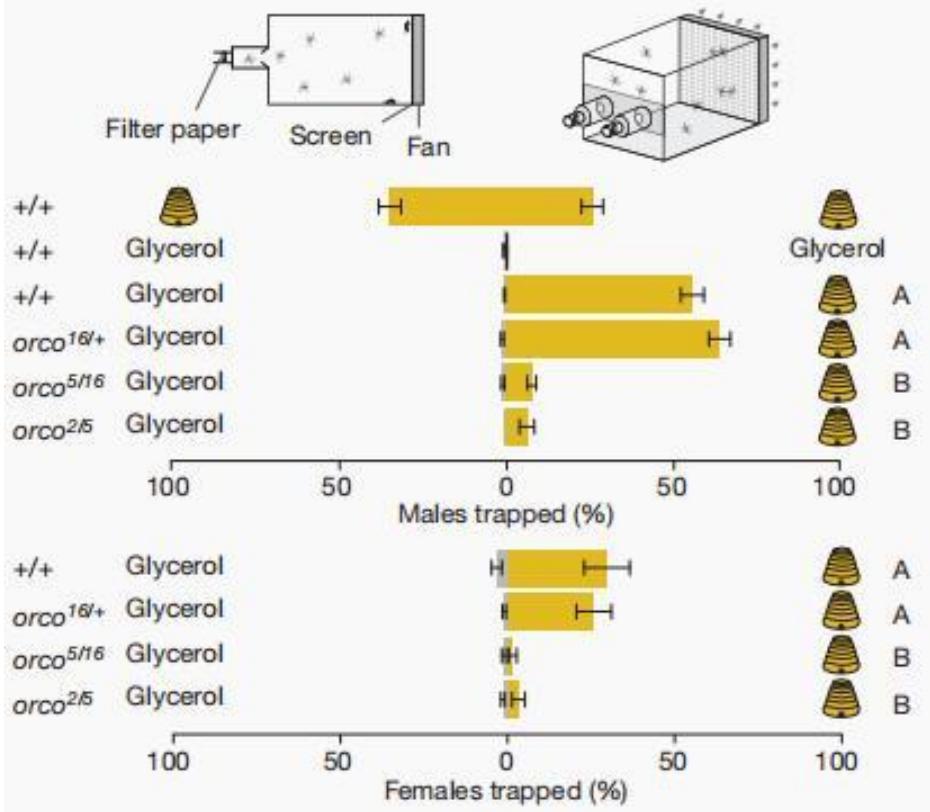
Multimodal floral cues guide mosquitoes to tansy inflorescences



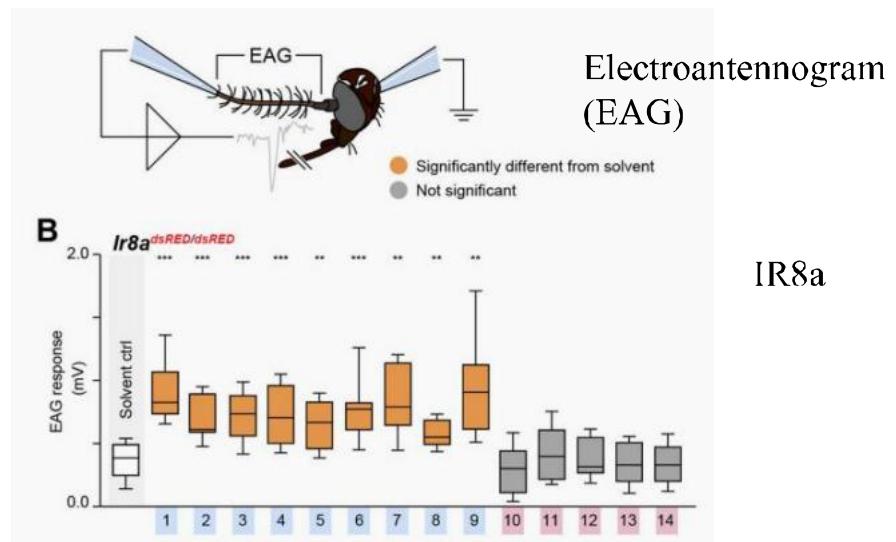
Peach DAH, Gries R, Zhai H, Young N, Gries G, Sci Rep, 2019

Mosquitoes Use Olfaction to Locate Nectar Sources

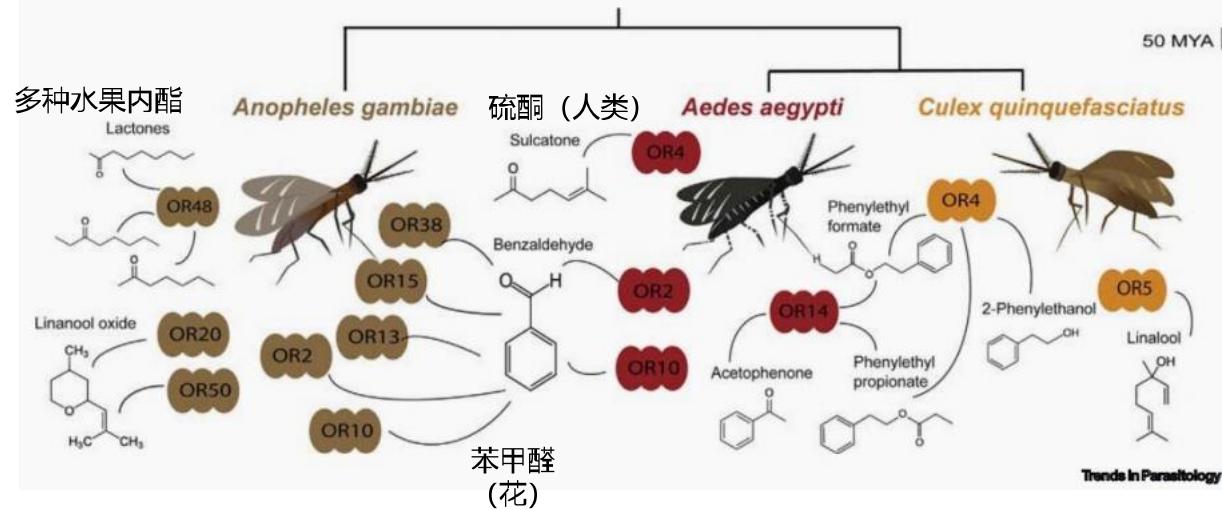
Disruption of honey odour detection in orco mutants.



DeGennaro M, McBride CS, et al., *Nature*, 2013



Raji JI, Melo N, Castillo JS, Gonzalez S, *Curr Biol*, 2019



芳樟醇氧化物

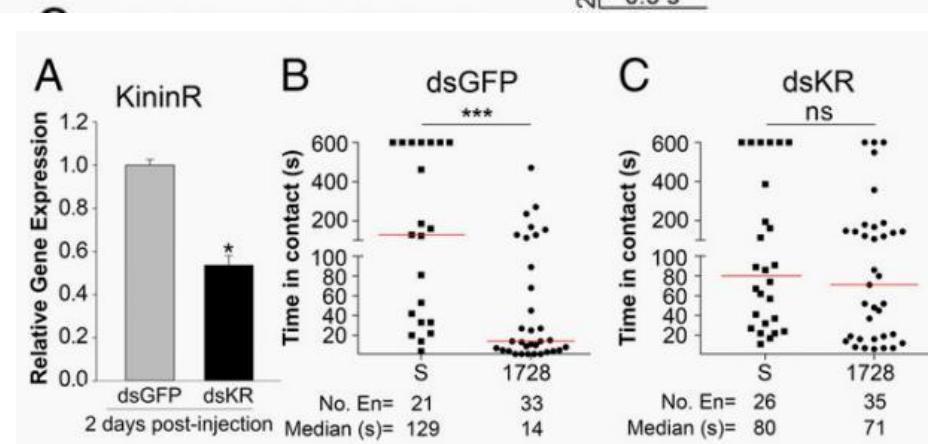
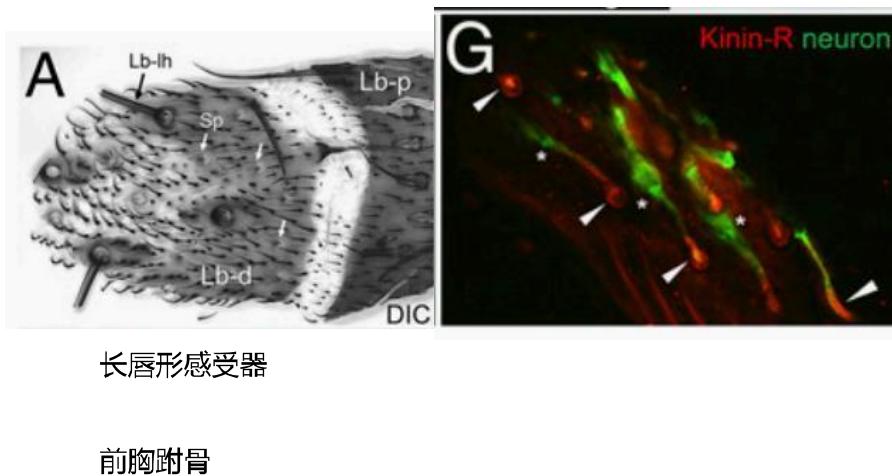
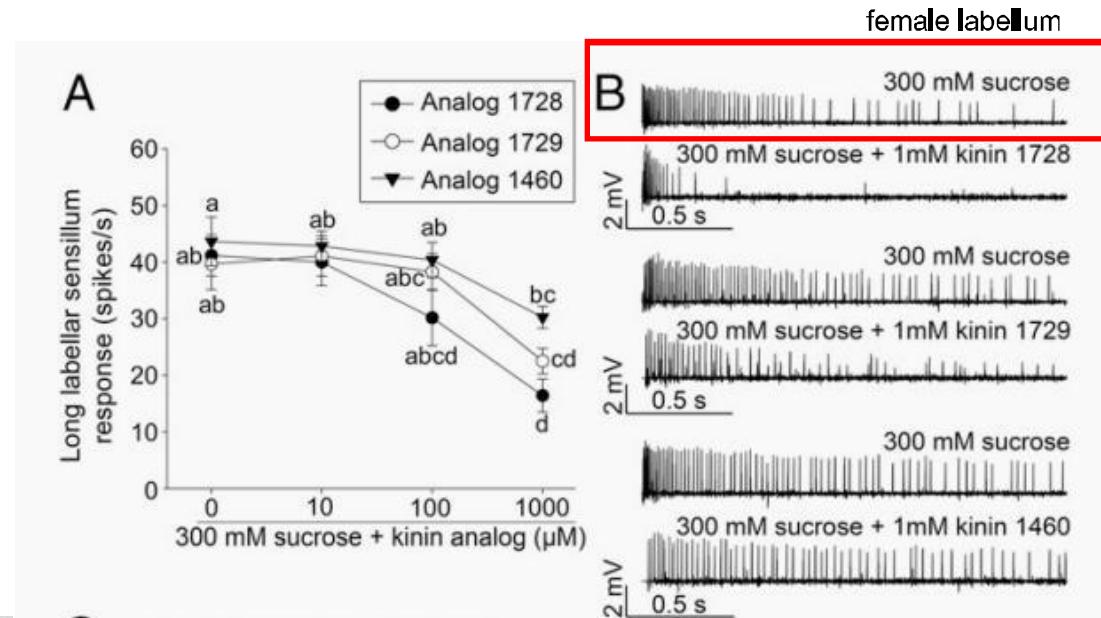
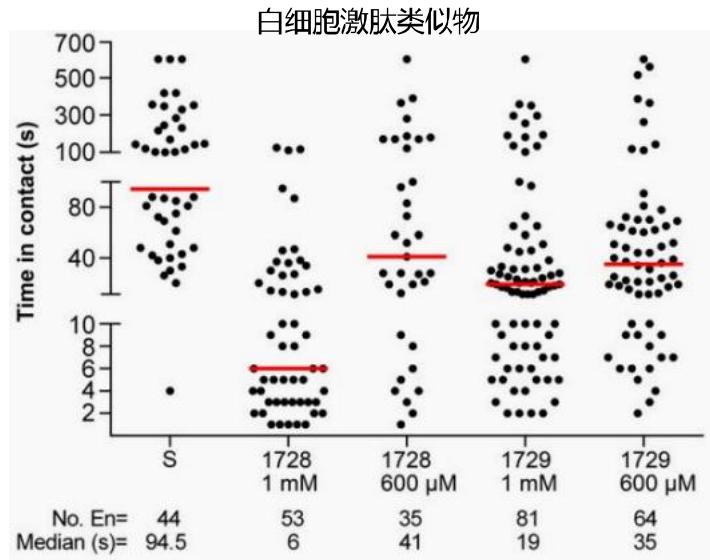
Barredo E, DeGennaro M., *Trends Parasitol*, 2020

Leucokinin mimetic elicits aversive behavior in *Aedes aegypti* and inhibits the sugar taste neuron

Movie S1

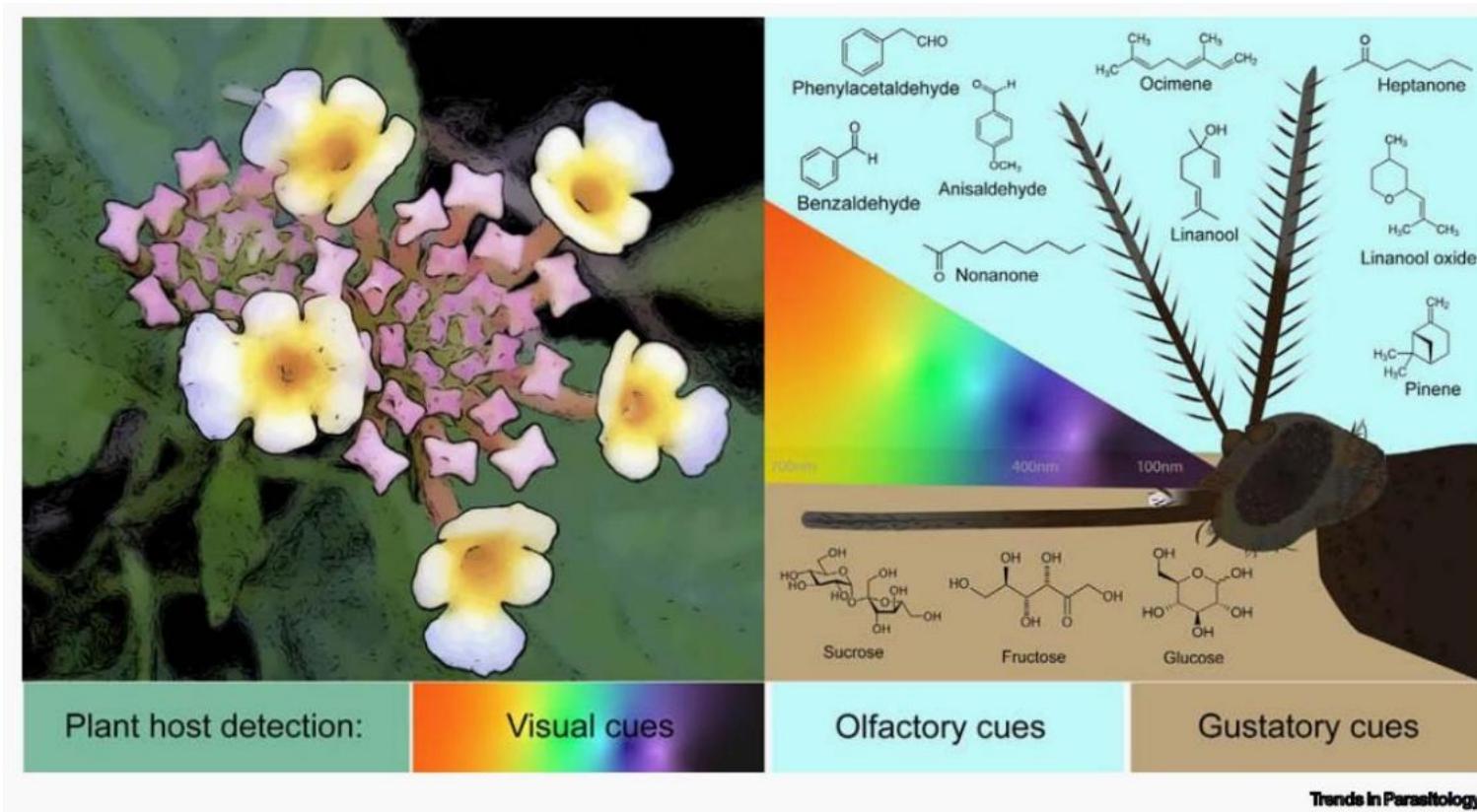
Females of *A. aegypti* feeding on
kinin analog 1728 at 1 mM
(10% sucrose solution, 0.1% Evans blue)

Leucokinin mimetic elicits aversive behavior in *Aedes aegypti* and inhibits the sugar taste neuron



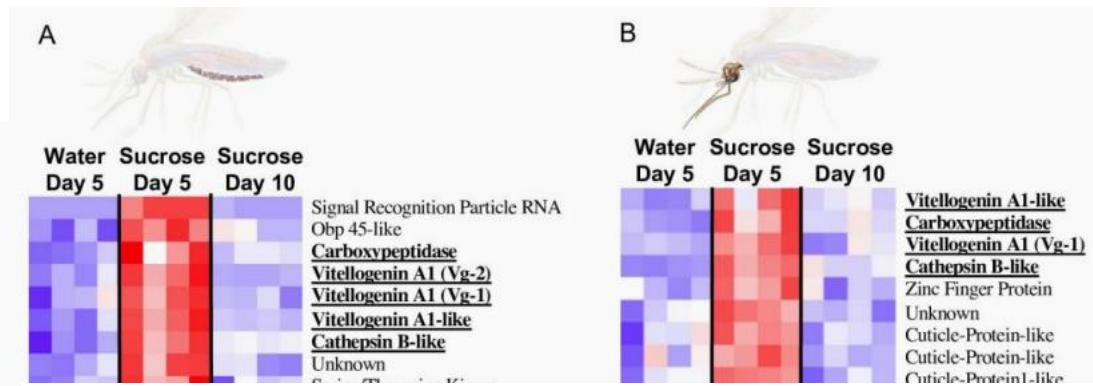
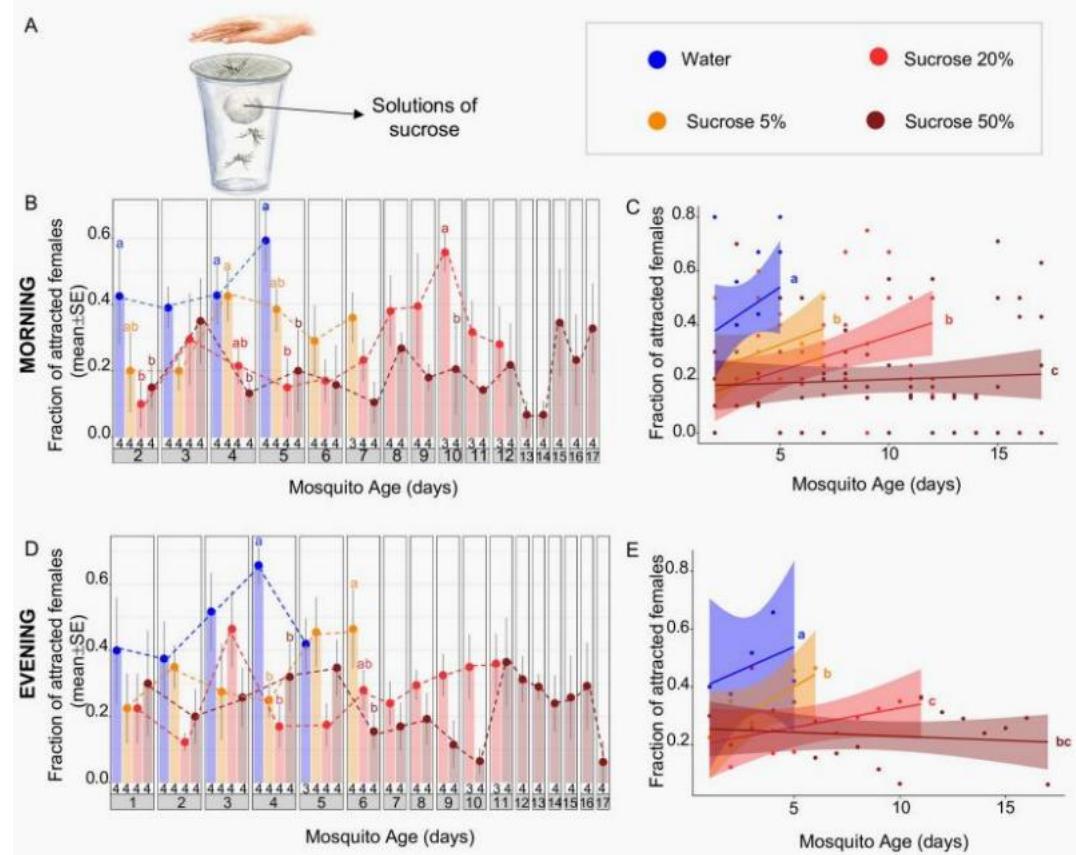
Kwon H, Ali Agha M,
et al., PNAS. 2016

Not Just from Blood: Mosquito Nutrient Acquisition from Nectar Sources

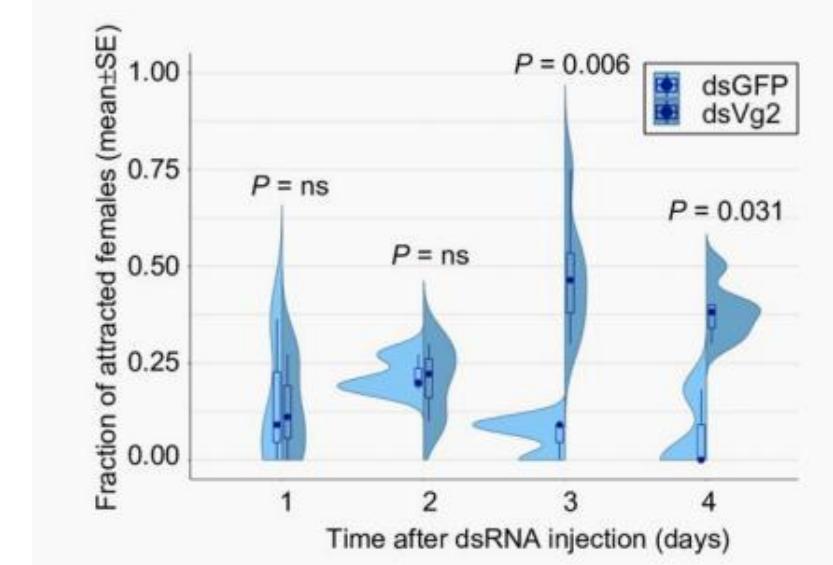


Barredo E, DeGennaro M., *Trends Parasitol*, 2020

Sugar feeding reduces host-seeking behaviour



Vg-2 expression regulates host-seeking behaviour

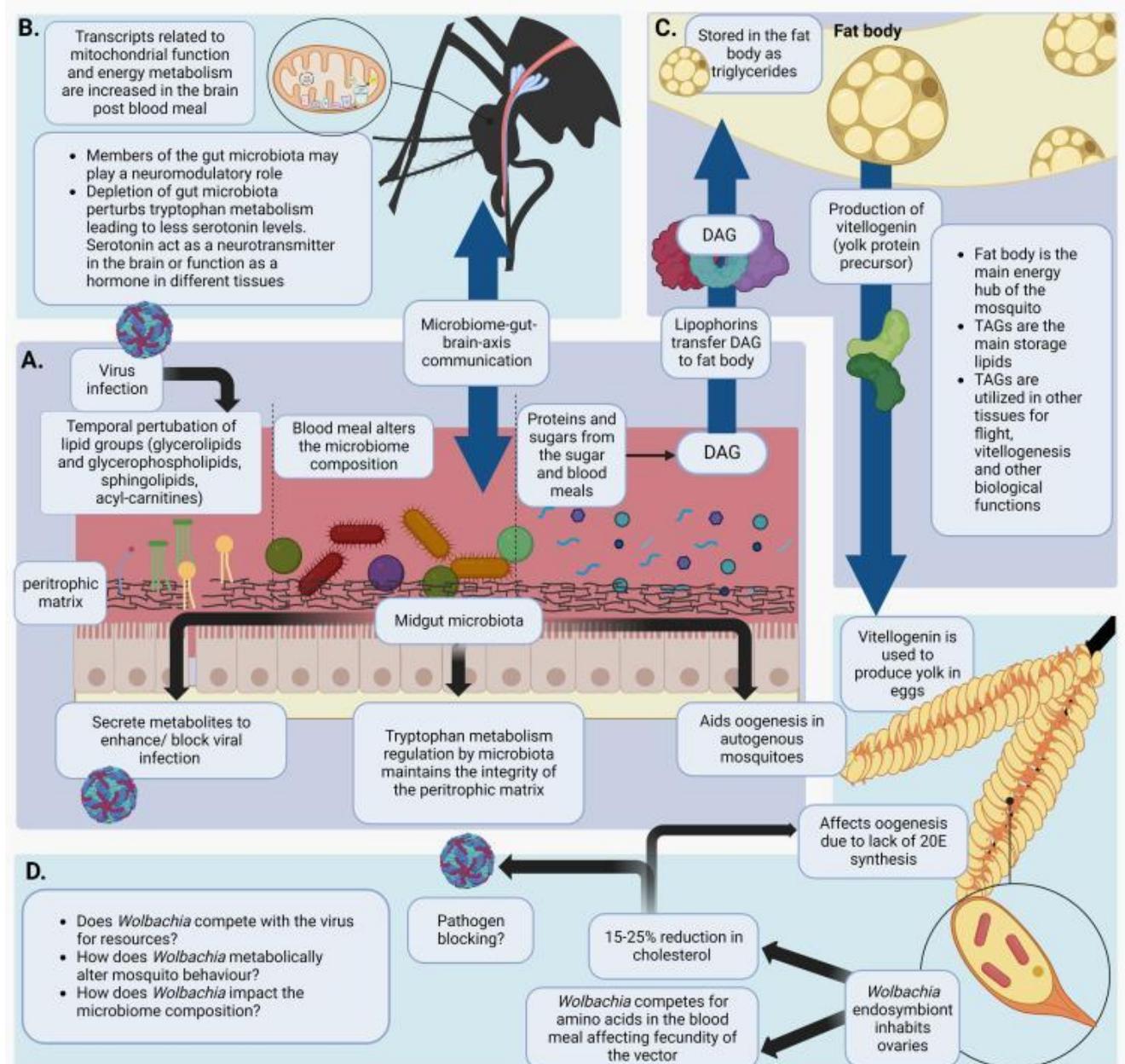


Why blood feed?

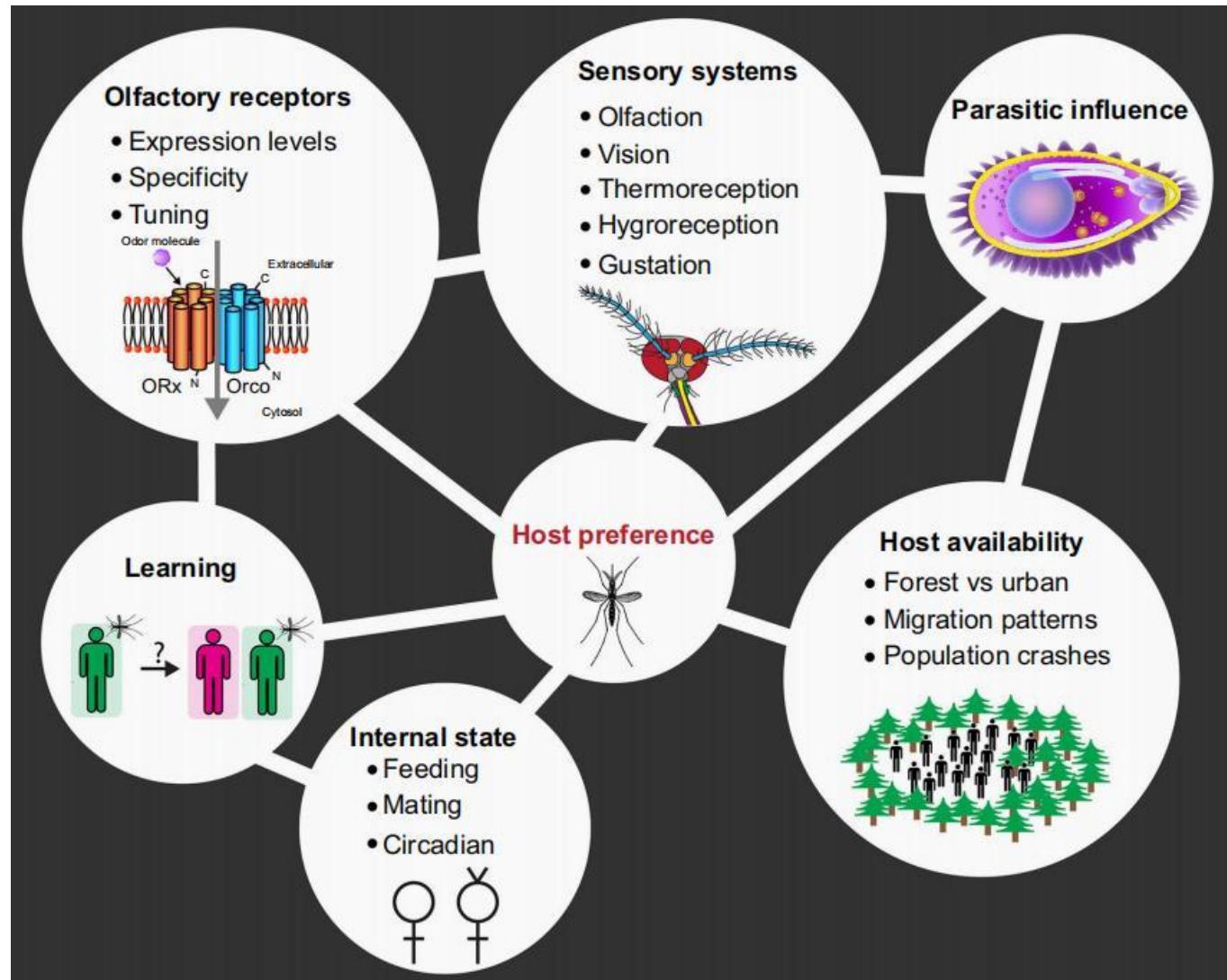
Blood meal induced metabolic changes in the mosquito.



Ratnayake OC, Chotiwon N, et al.,
Front Cell Infect Microbiol, 2023



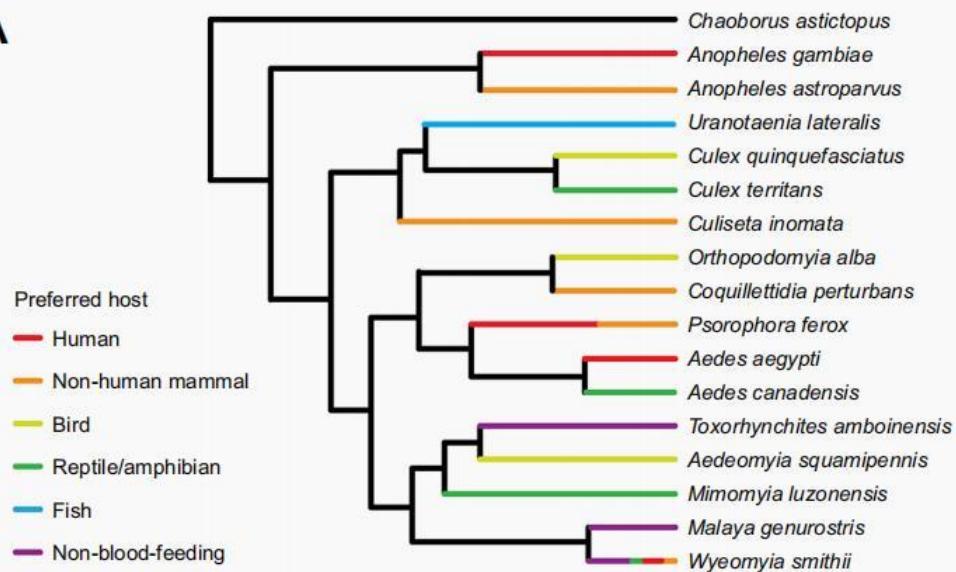
Factors influencing mosquito host-preference behaviors



Wolff GH, Riffell JA., *J Exp Biol*, 2018

Host preferences in specialists and generalists

A



anthropophagic: feed often on humans

ornithophilic: prefer birds to mammals

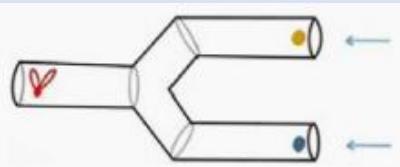
mammalophilic: prefer mammals

zoophilic: often on non-human animals

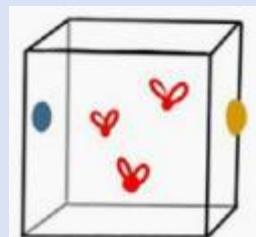
Specialists & Generalists & Opportunistic

Methods to determine host preference: behavioral observation and blood meal analyses

Behavioral observation



Y-tube



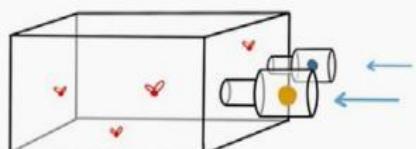
Choice chamber



Wind tunnel



Baited traps



Dual-port olfactometer

One limitation:

Certain genetic traits may get lost.

The preference may also be lost.

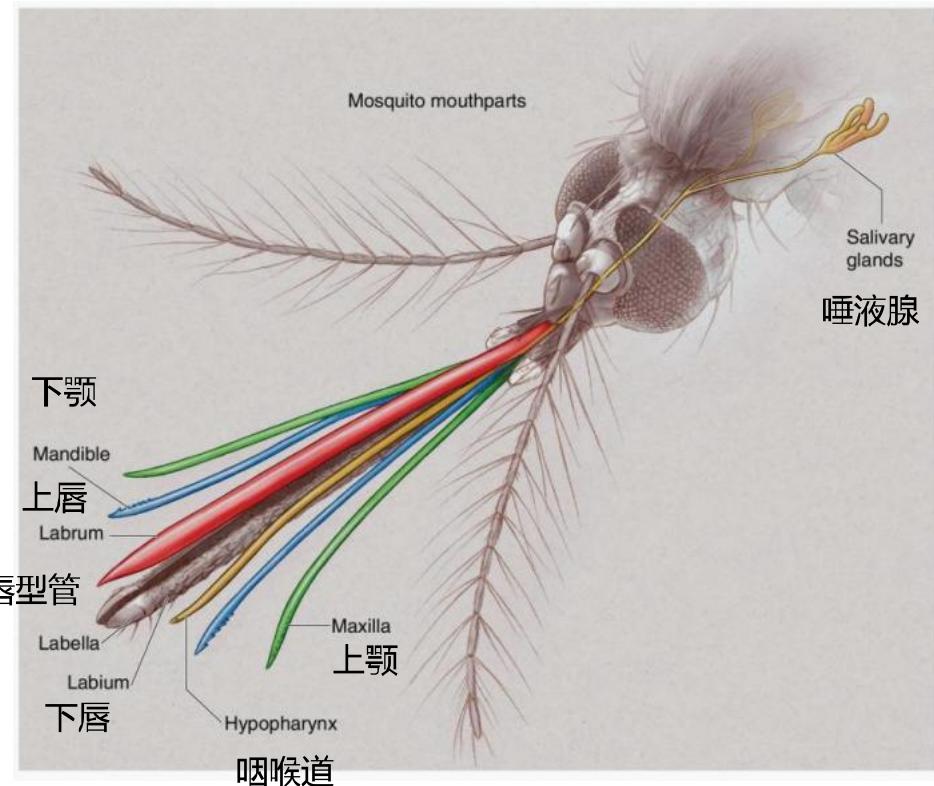
Blood meal analyses

The origin of blood meals can be assessed by:
multiplex PCR
microsphere assay
microsatellites
enzyme-linked immunosorbent assay (ELISA)
precipitin test.

One limitation:

May be favor of the most abundant
host species locally available.

Blood feeding phases



Penetration Time

insertion of the stylets into the host skin and usually lasts <10 sec

Probing Time

involves saliva secretion into the skin

Some researchers group penetration and probing time as the exploratory phase for blood.

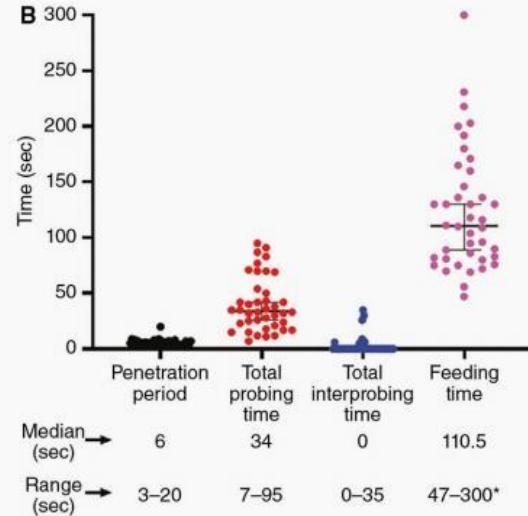
Feeding Time

an active phase of blood ingestion and engorgement

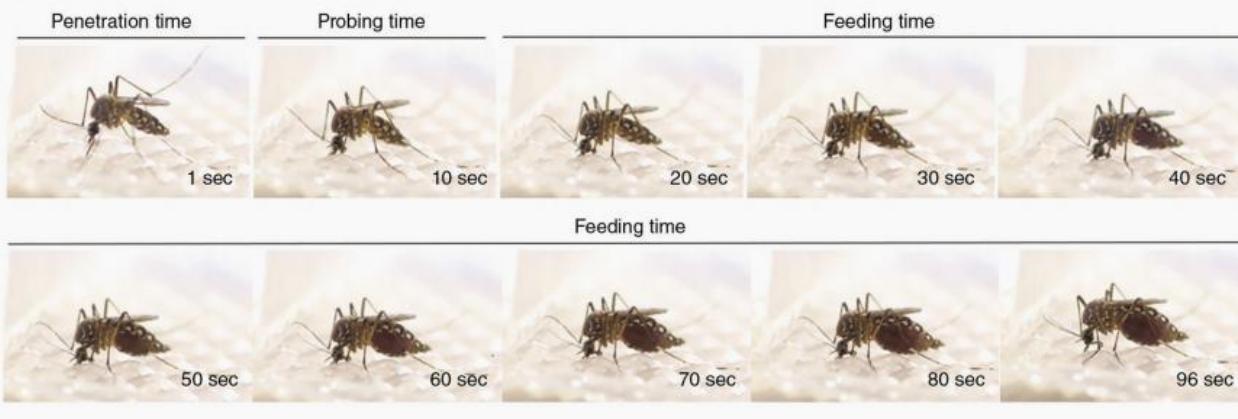


Evaluation of Aedes aegypti Penetration, Probing, and Feeding Times on Mice.

A



C



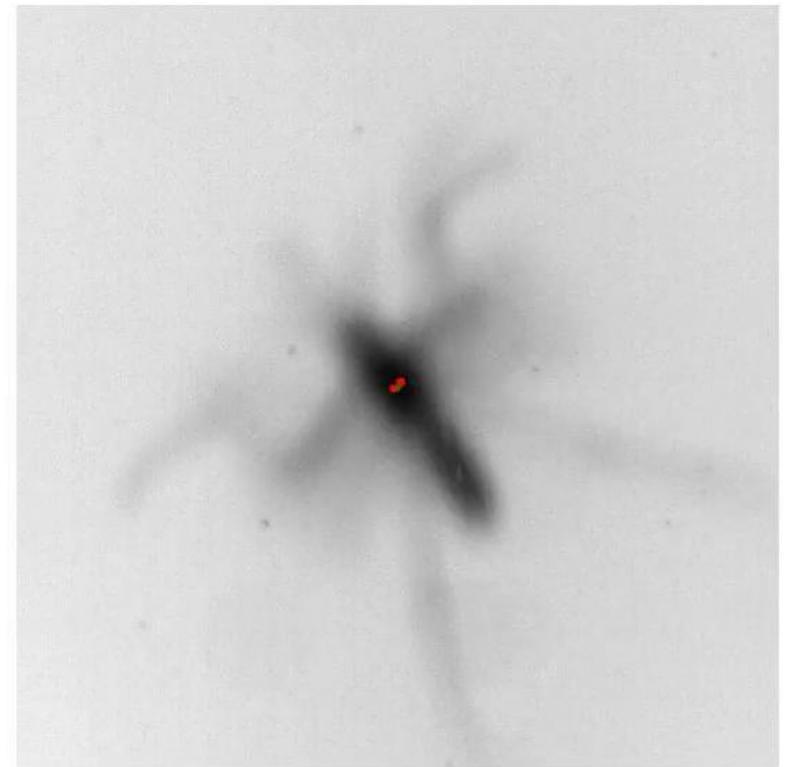
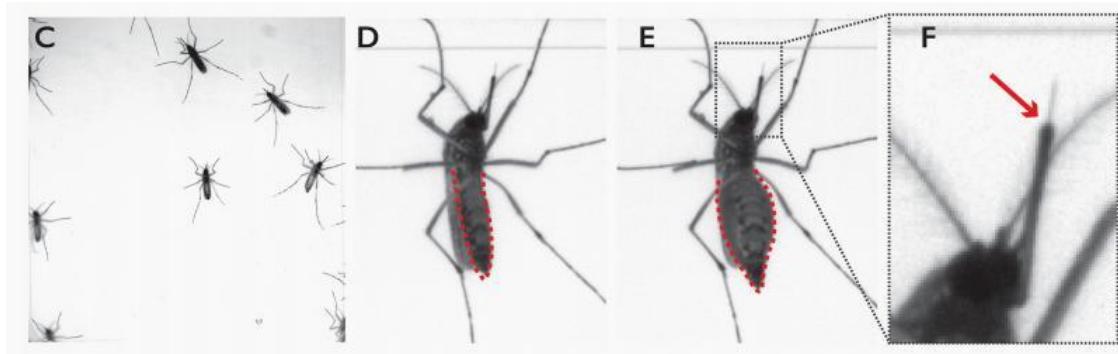
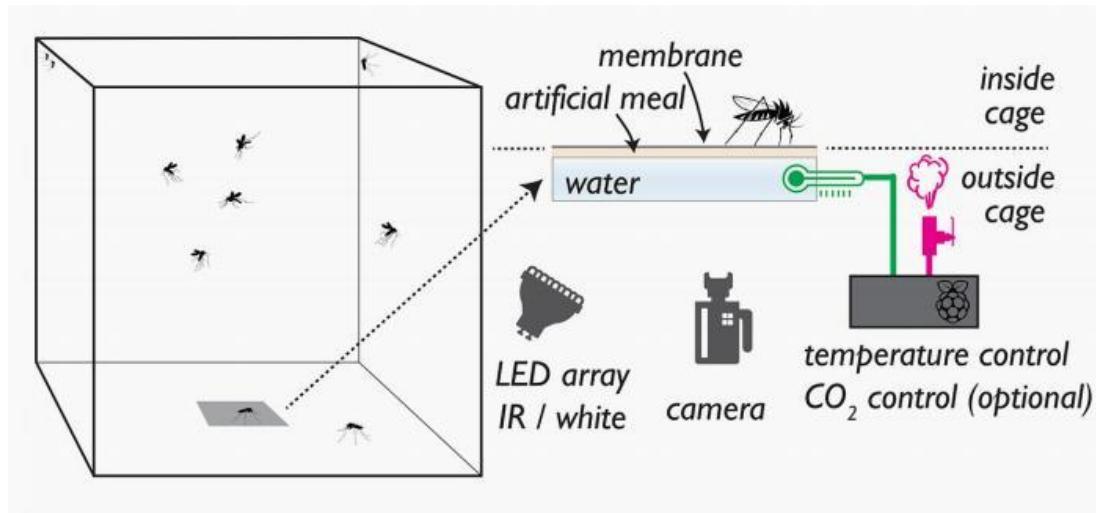
The time between probing times is called interprobing time.

Not interested in blood feeding

Solution:

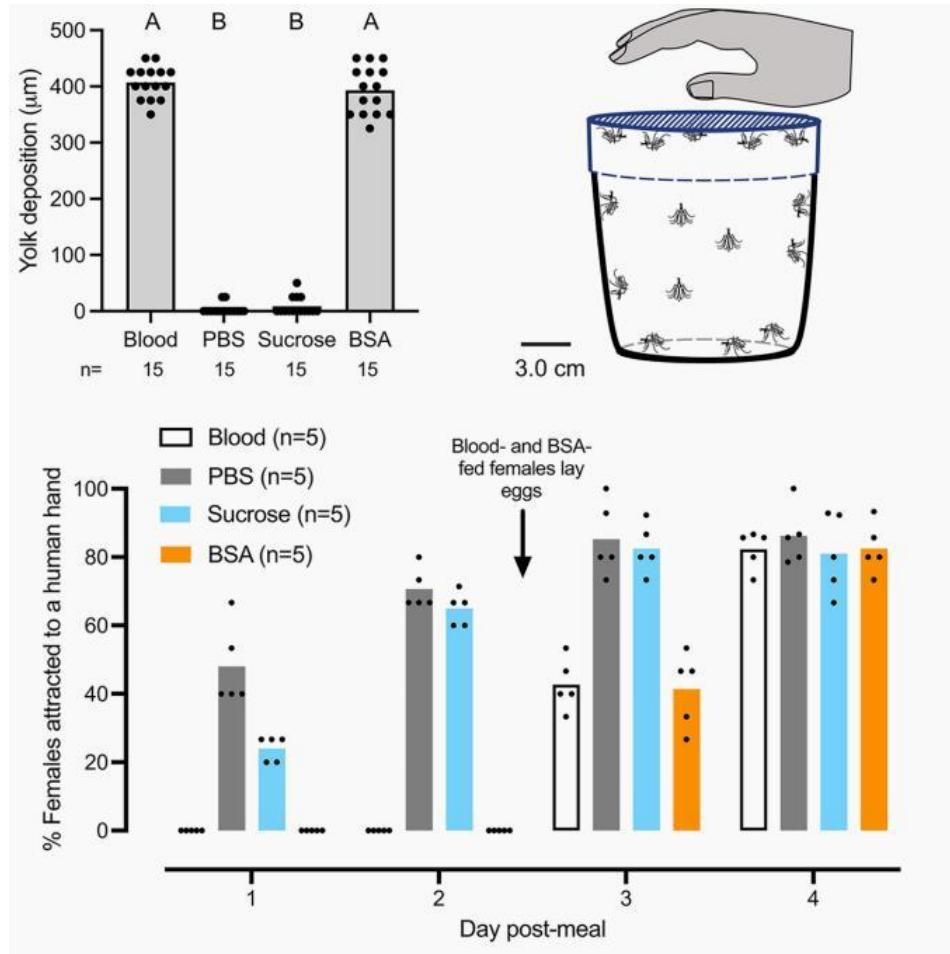
- Ensure that the temperature of the mouse did not drop .Perform in a room at 24°C– 26°C.
- Breathing near the mosquito vial
- Replace sugar cotton balls with water cotton balls in 2 d before the experiment. The night before the experiment, remove the water cotton balls.

BiteOsce, an open platform to study mosquito biting behavior

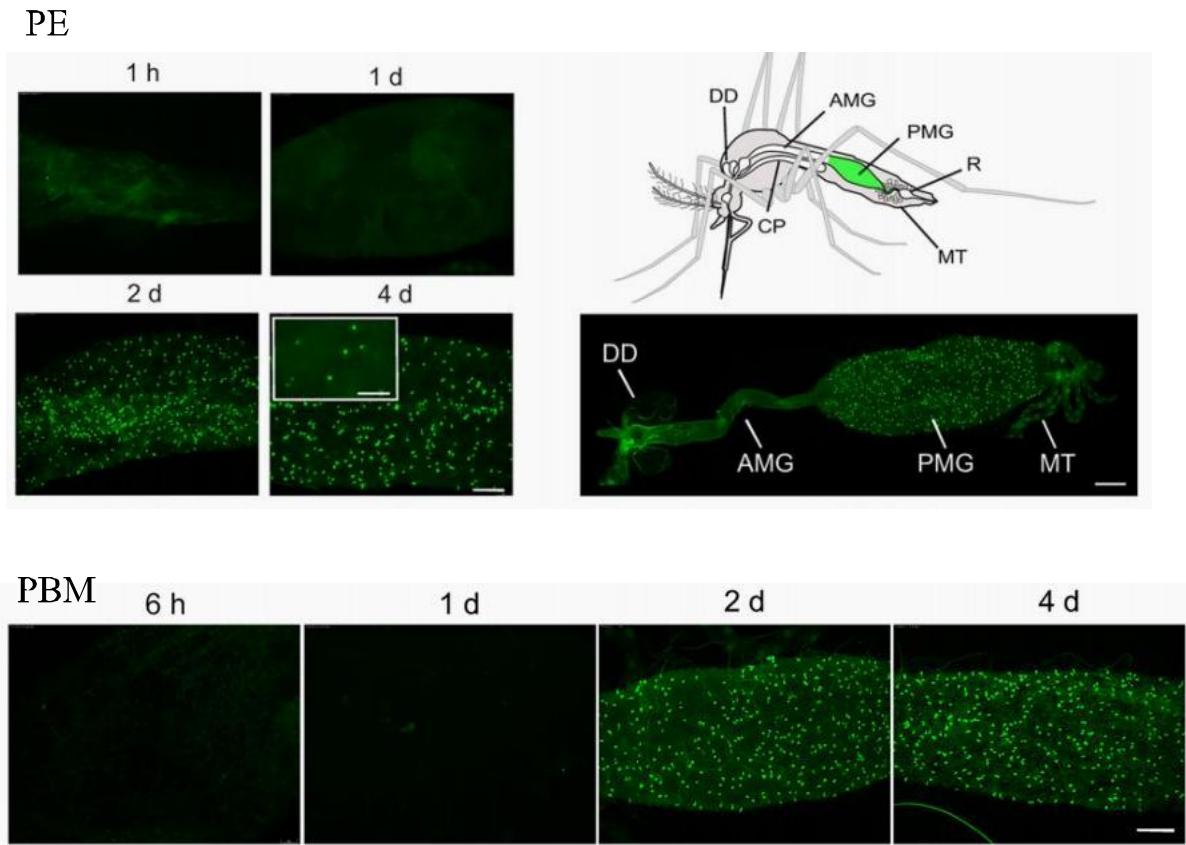


Hol FJ, Lambrechts L, Prakash M., *Elife*, 2020

Neuropeptide F and RYamide regulate host attraction in the mosquito *Aedes aegypti*



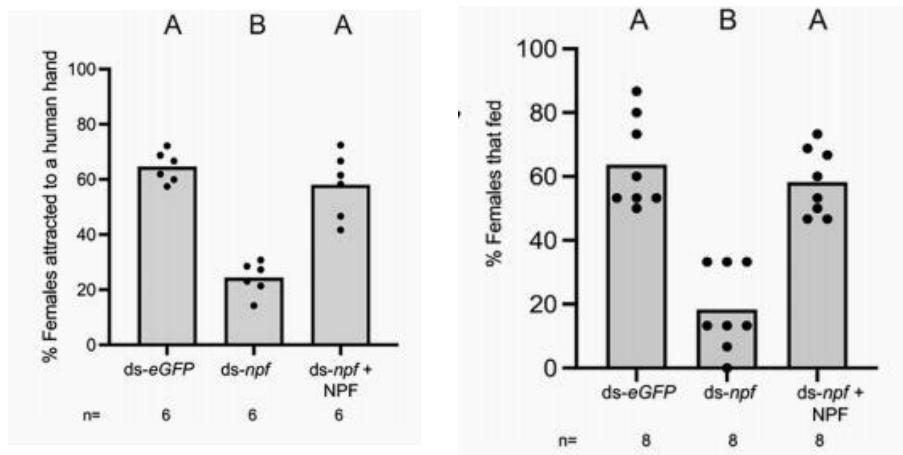
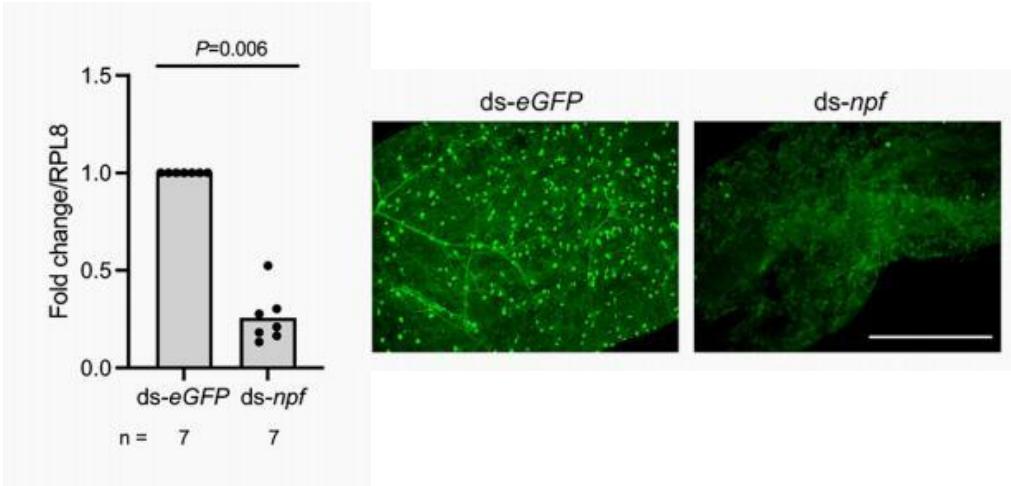
Only blood and protein meals activate the vitellogenic phase and suppress host attraction.



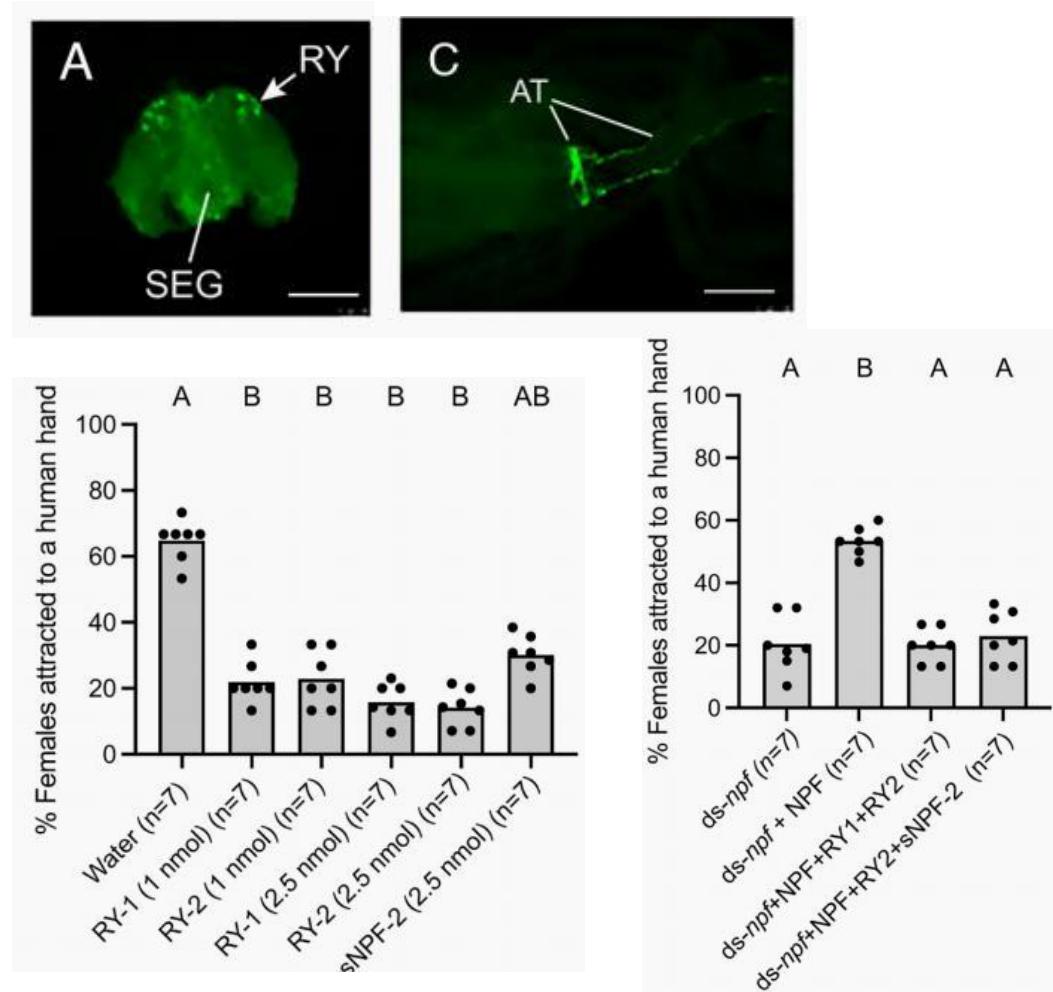
NPF accumulates in EECs of the posterior midgut after adult emergence but rapidly depletes after blood feeding.

Dou X, Chen K, Brown MR, Strand MR. *PNAS*, 2024

Neuropeptide F and RYamide regulate host attraction in the mosquito *Aedes aegypti*



RNAi knockdown of midgut NPF reduces host attraction and blood feeding.



RY-amides suppress host seeking.

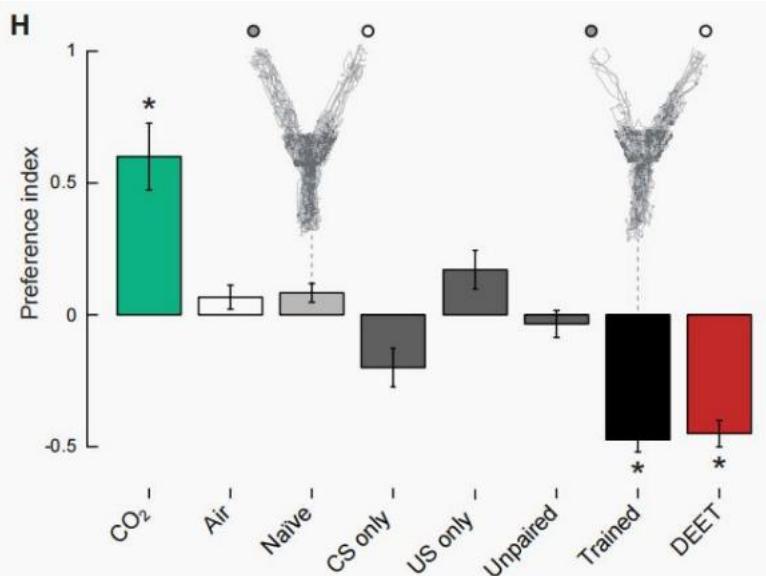
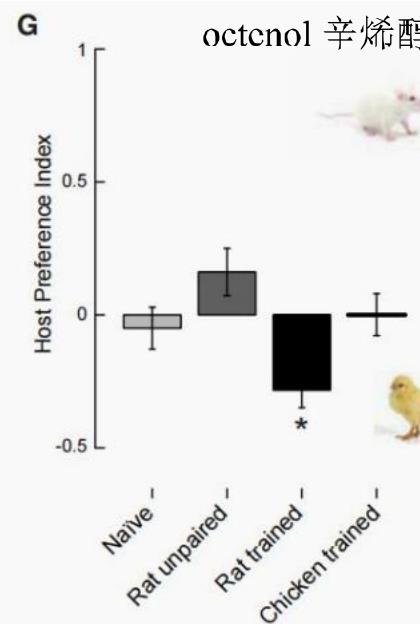
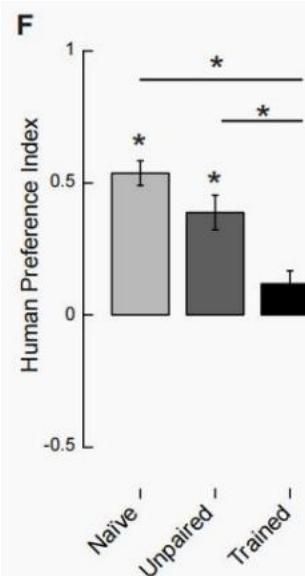
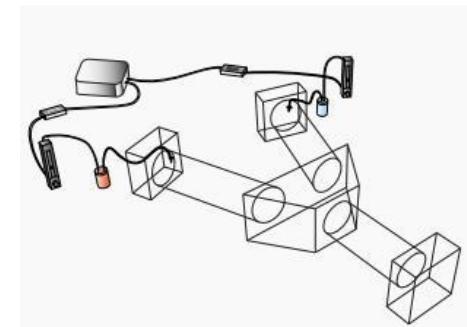
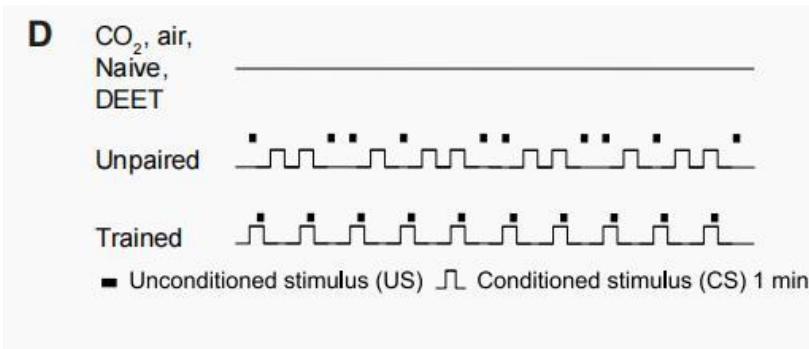
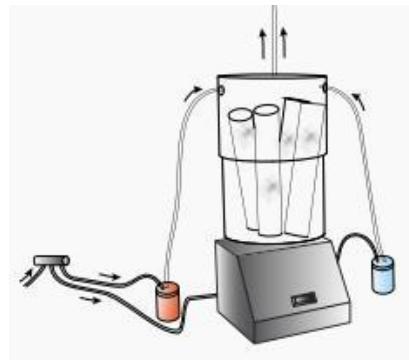
Dou X, Chen K, Brown MR, Strand MR., PNAS, 2024

Can host preferences be changed?

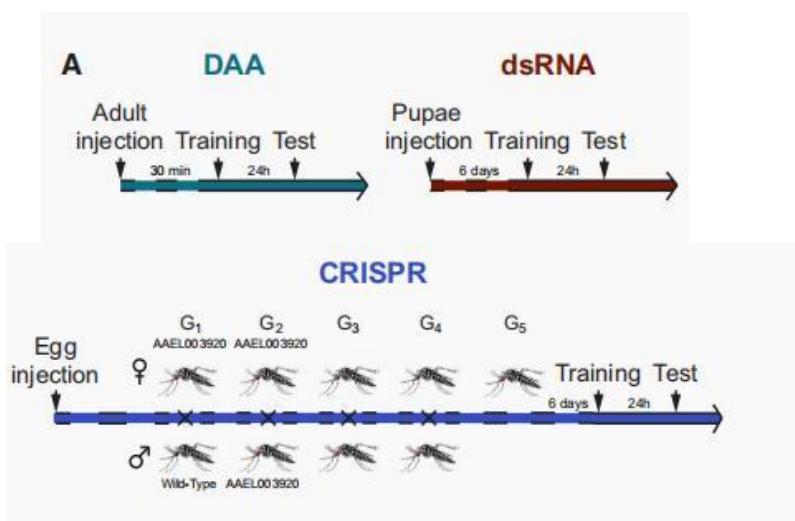
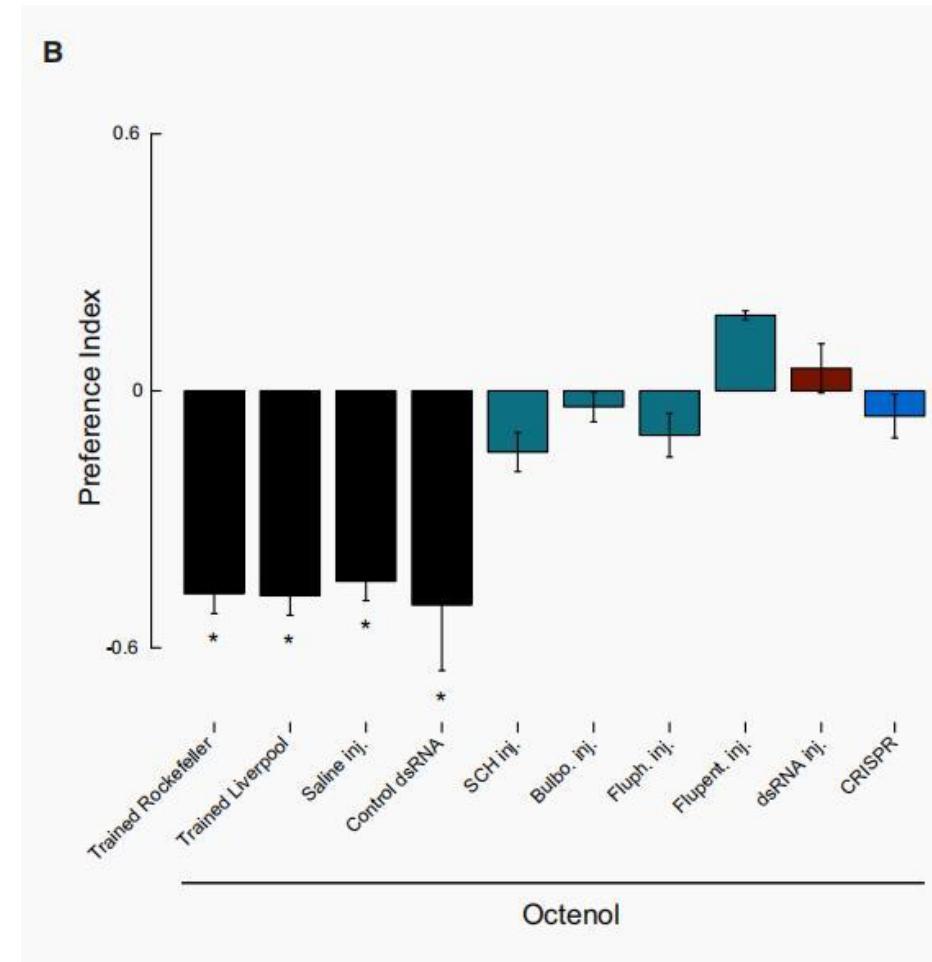
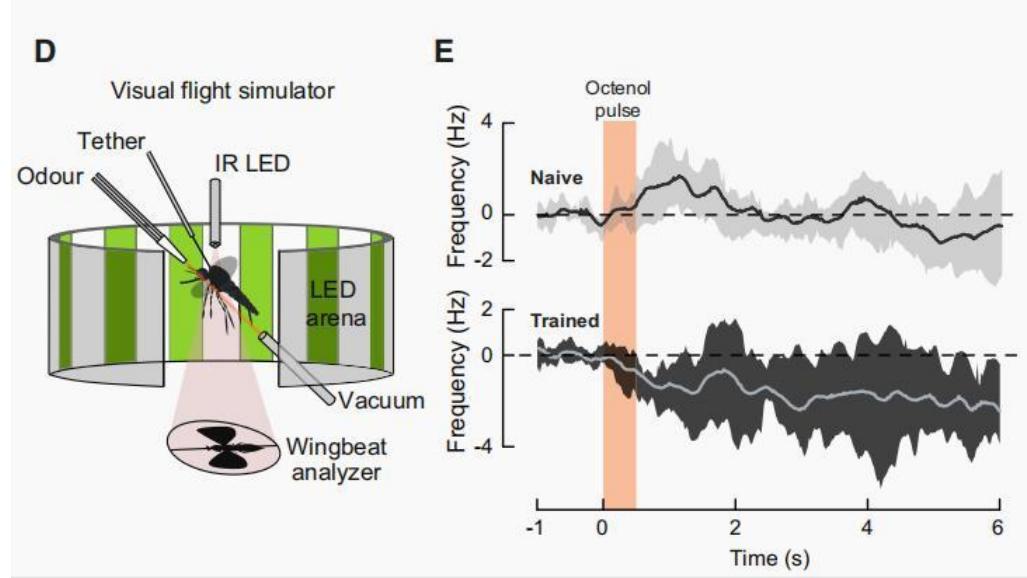
- Learning
- Viral infection

Learning:

Mosquitoes Aversively Learn Host Odors and Single Odorants

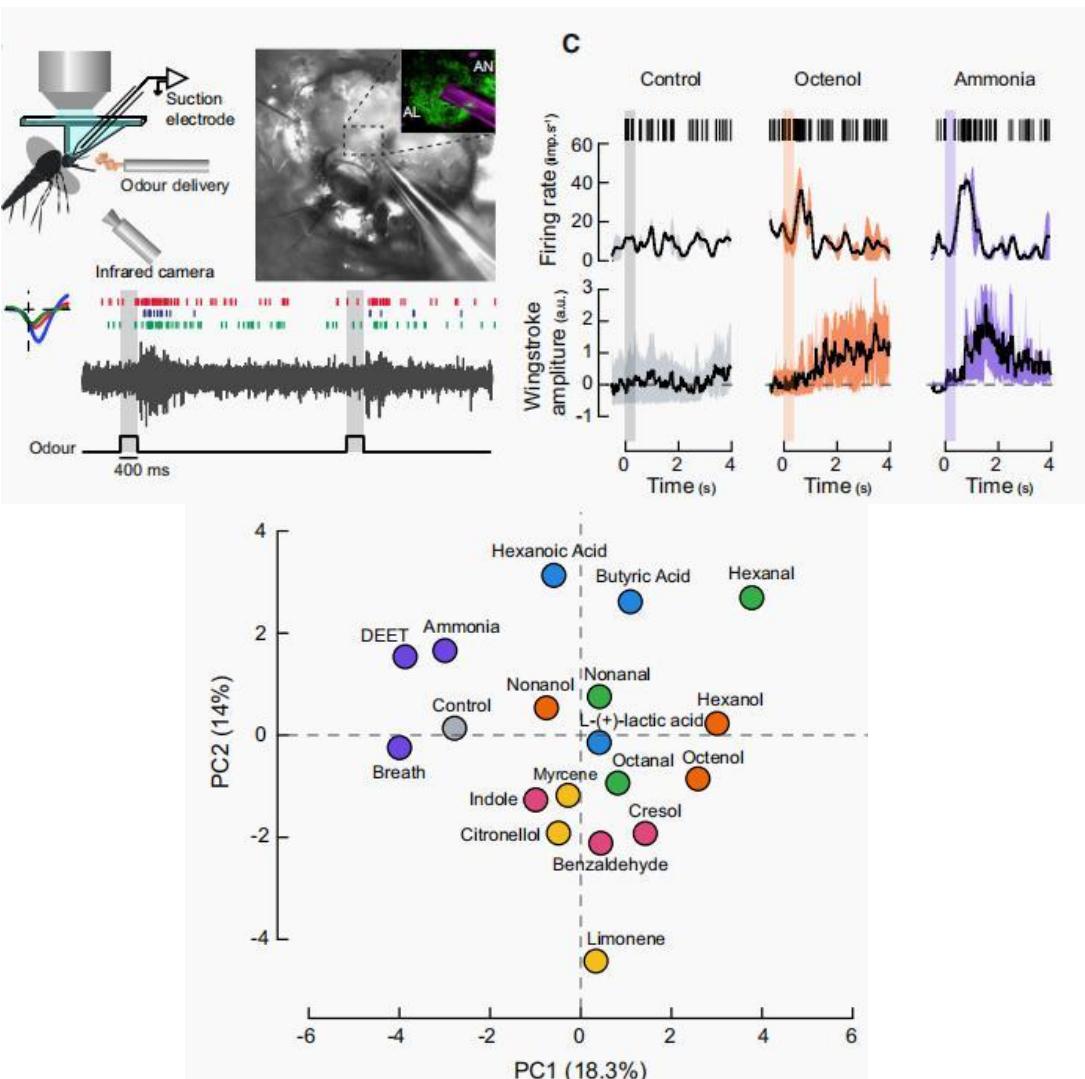


Dopamine Is Critical for Aversive Learning

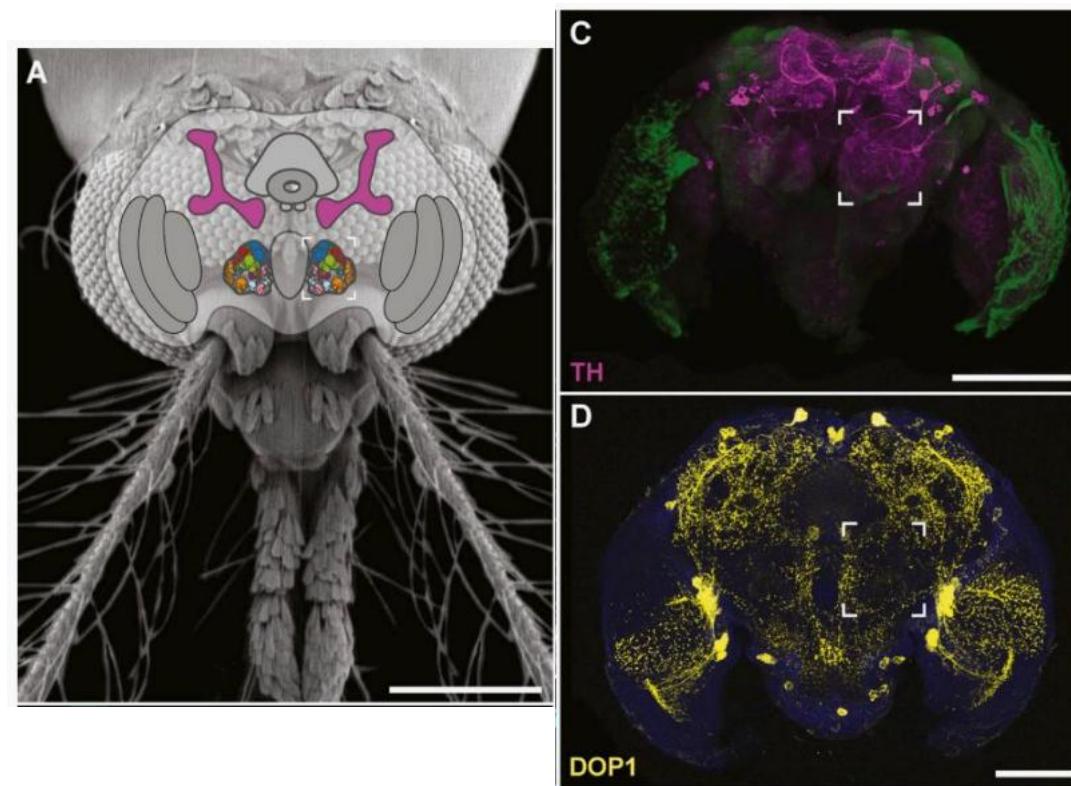


Vinauger C, Lahondère C, et al., *Curr Biol*, 2018

Odor Stimuli Are Learned and Represented Differentially in the Mosquito Brain

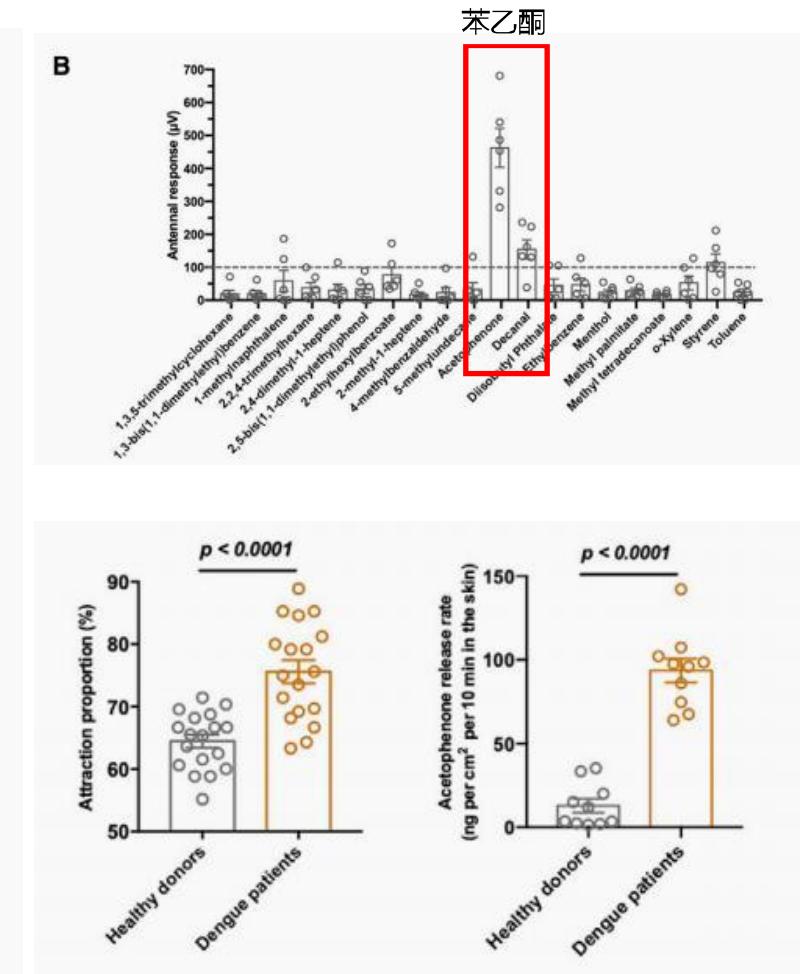
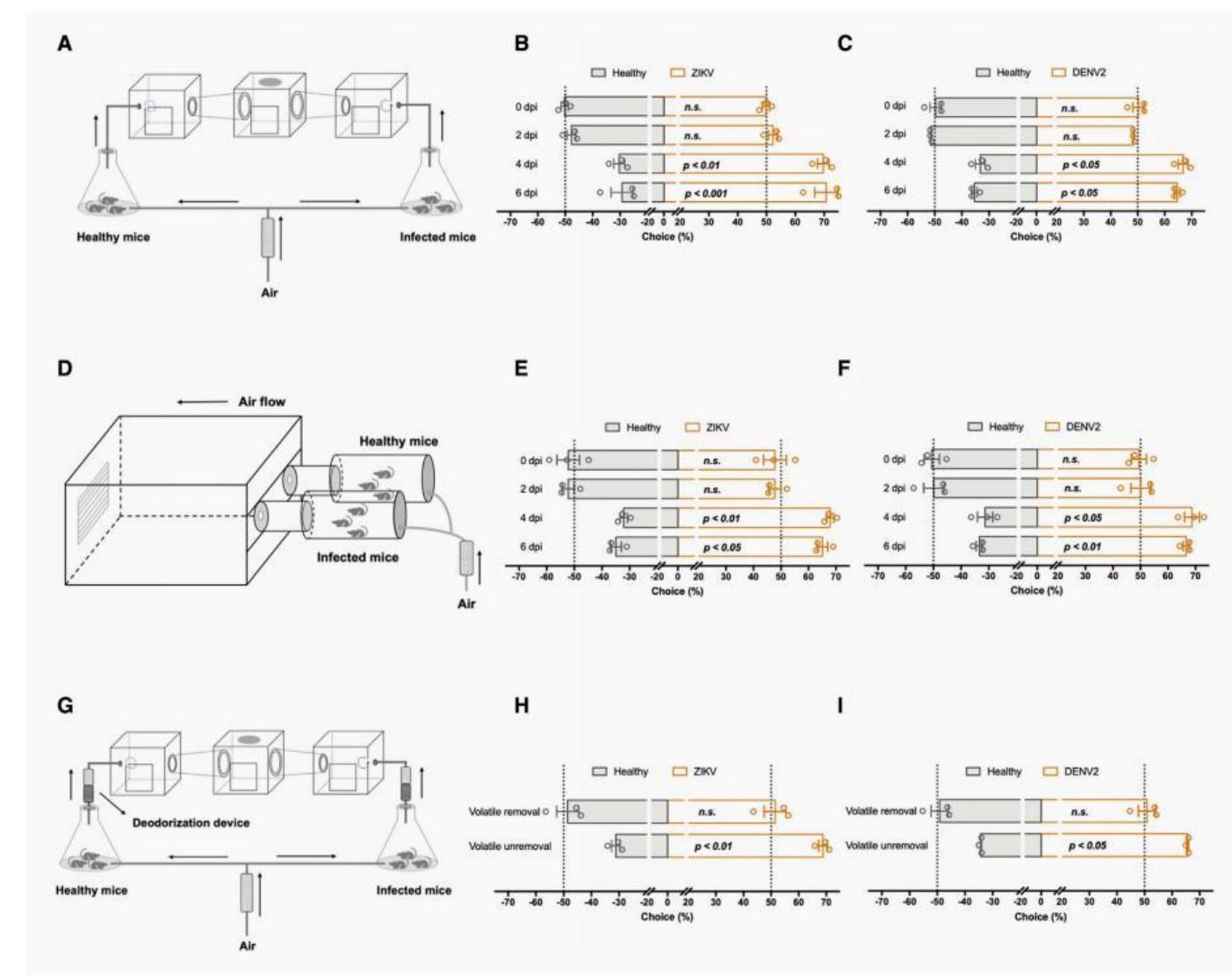


Tyrosine Hydroxylase and Dopamine Receptor Immunoreactivity



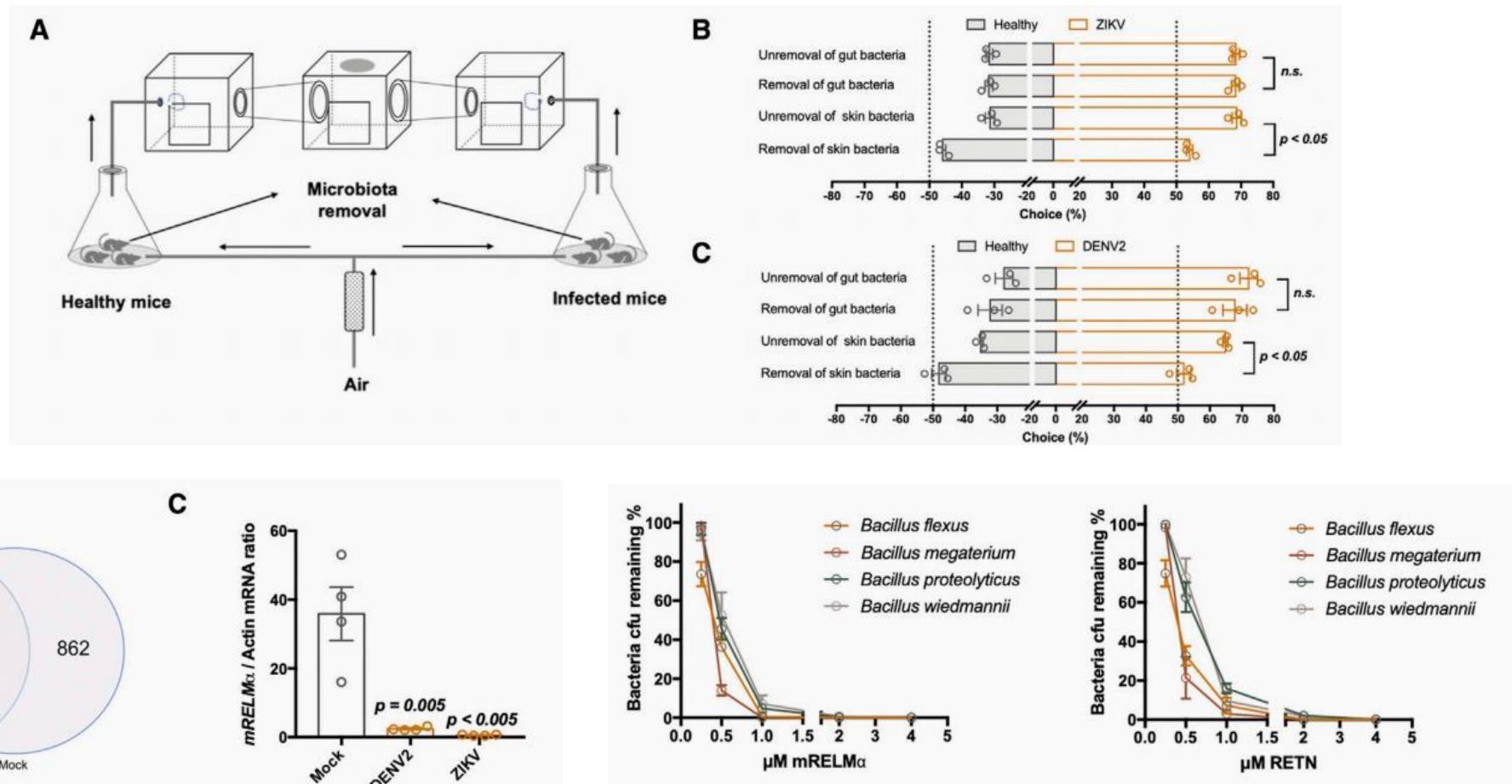
Vinauger C, Lahondère C, et al., *Curr Biol*, 2018

Viral infection: Mice infected by flaviviruses present a higher attraction to *A. aegypti*



Zhang H, Zhu Y, et al., *Cell*, 2022

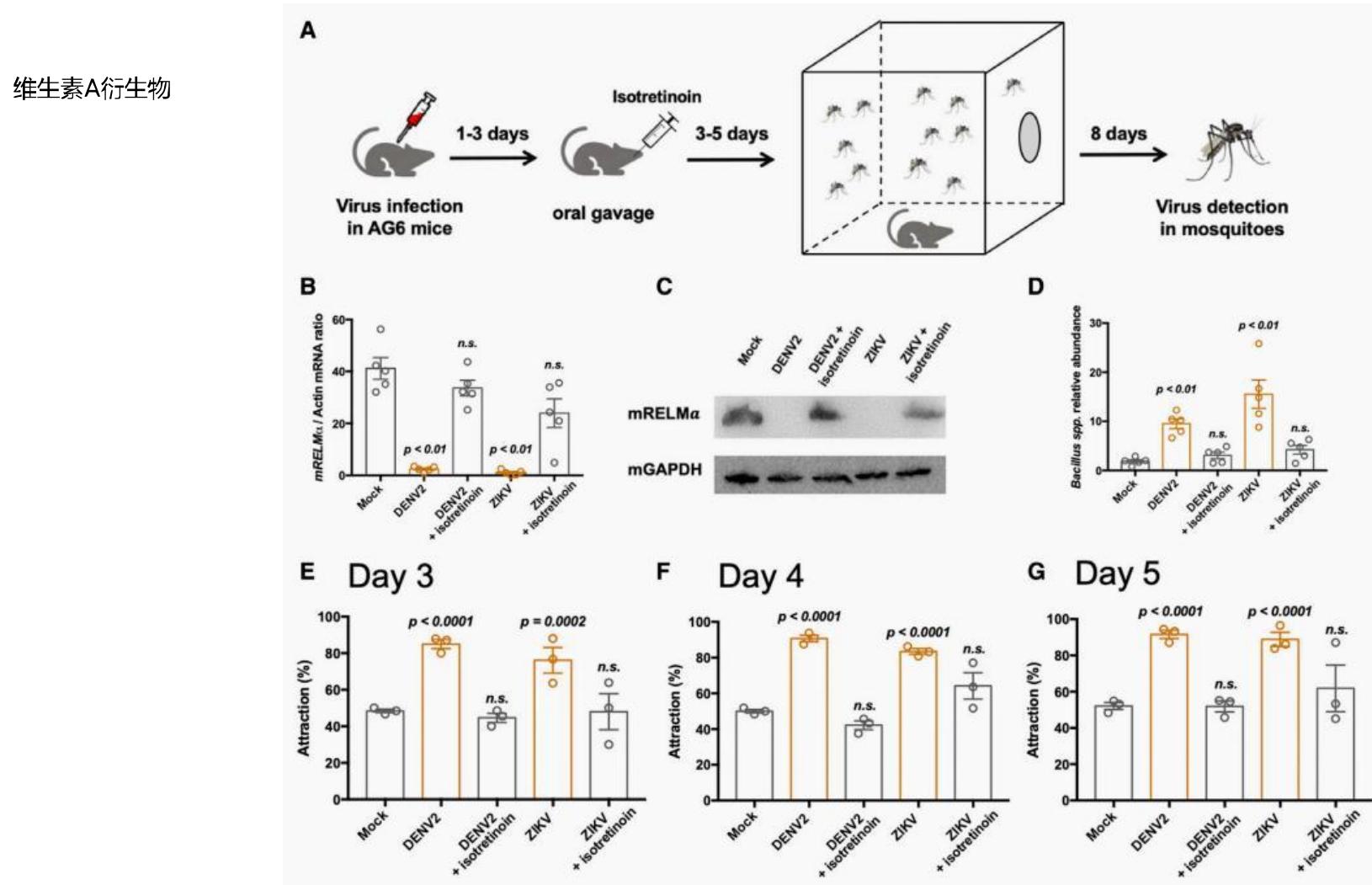
Skin microbiota regulated by flavivirus infection are responsible for releasing acetophenone for mosquito attraction



Retnla负责编码抵抗素样分子- α (RELM α)，一种抗菌蛋白

Zhang H, Zhu Y, et al., *Cell*, 2022

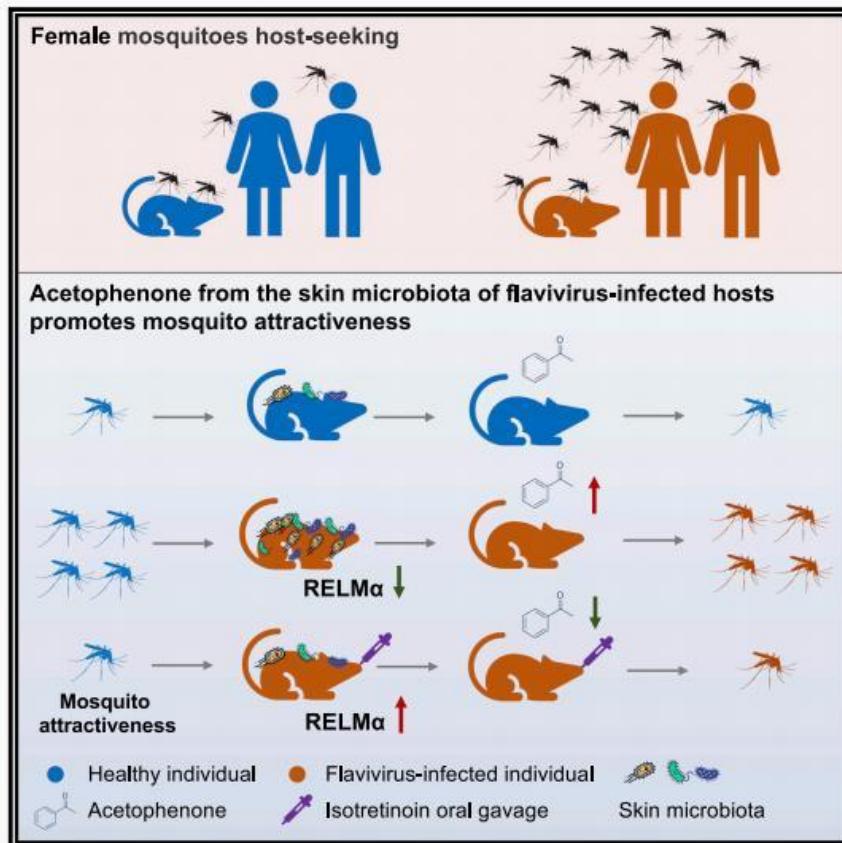
Dietary administration of a vitamin derivative to flavivirus-infected mice reduces the acetophenone cue to mosquitoes and viral transmission



Zhang H, Zhu Y, et al., *Cell*, 2022

A volatile from the skin microbiota of flavivirus-infected hosts promotes mosquito attractiveness

Graphical abstract



Authors

Hong Zhang, Yibin Zhu, Ziwen Liu, ..., Qiyong Liu, Penghua Wang, Gong Cheng

Correspondence

gongcheng@mail.tsinghua.edu.cn

In brief

Flaviviruses such as dengue and Zika modulate murine host skin bacterial communities to increase acetophenone-producing bacteria. Acetophenone is a mosquito attractant, and its increased production by flavivirus-infected humans and mice make them more attractive to mosquitoes, facilitating viral transmission by mosquito vectors.



技术专利

1. 程功, 肖小平。一种昆虫蛋白Hig及其编码基因和应用。授权号: ZL201410837328.9
2. 程功, 肖小平。获自埃及伊蚊的防御素及其编码基因和应用。授权号: ZL201310459570.2
3. 程功, 肖小平。获自埃及伊蚊的天蚕素及其编码基因和应用。授权号: ZL201310455321.6
4. 程功, 刘洋。一种用于登革热预防和/或治疗的疫苗。授权号: ZL201310181920.3。(申请PCT国际专利保护)
5. 程功, 刘建英, 刘洋。一种登革热双效疫苗的制备方法及其应用。授权号: ZL201510830549.3。(申请PCT国际专利保护)
6. 郑兆鑫, 严维耀, 程功, 刘明秋。一种口蹄疫基因工程多肽疫苗佐剂及其制备方法和应用。授权号: ZL200610147664.6。
7. 郑兆鑫, 赵欣, 程功, 严维耀, 刘明秋。猪干扰素基因家族及其作为抗猪蓝耳病药物及疫苗佐剂的应用。授权号: ZL201010208935.0。
8. 程功, 杜森焱。ZIKV-NS1蛋白及其在制备寨卡病毒传播阻断疫苗中的应用。申请号: 201711327739.3。
9. 程功, 朱毅斌。调控GABAA受体的物质在调控昆虫虫媒病毒载量中的应用。申请号: 201710803946.5。
10. 程功, 太万博, 冯胜勇, 庞慕加。一种靶向激活体液免疫和细胞免疫的核酸疫苗载体构建及应用。申请号: 202110655890.2。

从事蚊媒病毒感染传播机制与抗病毒免疫研究。研究主要涉及蚊媒病毒在哺乳动物宿主及节肢动物媒介中的感染机制，希望通过多种实验手段鉴定出与病毒感染高度相关的宿主蛋白，并利用鉴定出的宿主蛋白作为干预靶点，阻断虫媒病毒在自然界传播。

- **登革热病毒研究**

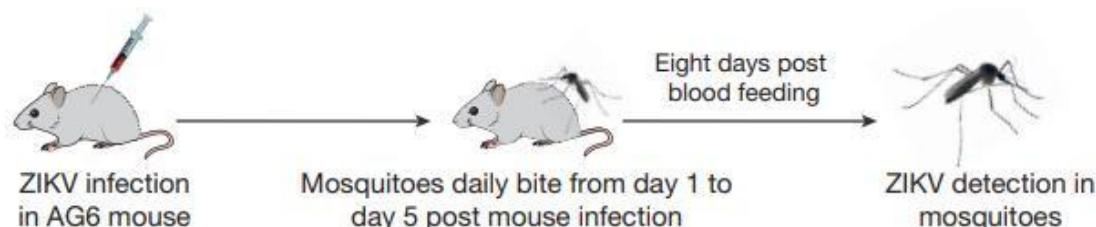
首次发现了登革病毒非结构蛋白NS1，回答了“登革病毒如何从宿主传播到蚊虫”这一重要科学问题。

- **首次阐明寨卡病毒暴发机制**

首先发现寨卡病毒的NS1蛋白同样也具有辅助病毒感染蚊虫的功能

- **疫苗研发**

新型mRNA疫苗和纳米颗粒疫苗



•代表性论著

1. Zhang H*, Zhu YB*, Liu ZW, Peng YM, Peng WY, Tong LQ, Wang JL, Liu QY, Wang PH, Cheng G#. (2022) A volatile from the skin microbiota of flavivirus-infected hosts promotes mosquito attractiveness. **Cell.** In Press.
2. Tong LQ*, Xiao XP*, Li M*, Fang SS*, Yu X, Zhu YB, Wu CL, Tian DY, Yang F, Sun J, Qu J, Zheng NZ, Liao SM, Tai WB, Feng SY, Zhang LM, Li YH, Wang L, Han XL, Sun SH, Yang L, Zhao JC, Liu WJ, Liu XH, Wang PH, Li L#, Zhao GY#, Zhang RL#, Cheng G#. (2022) A glucose-like metabolite deficient in diabetes inhibits cellular entry of SARS-CoV-2. **Nature Metabolism.** 4: 547–558.
3. Yu X, Shan C, Zhu YB, Ma EH, Wang JL, Wang PH, Shi PY, Cheng G#. (2021) A mutation-mediated evolutionary adaptation of Zika virus in mosquito and mammalian host. **Proceedings of the National Academy of the Sciences.** 118: e2113015118.
4. Zhu YB, Tong LQ, Nie KX, Wiwatanaratanabutr I, Sun P, Li QQ, Yu X, Wu P, Wu TS, Yu C, Liu QY, Bian ZQ, Wang PH, Cheng G#. (2019) Host serum iron modulates dengue virus acquisition by mosquitoes. **Nature Microbiology.** 4: 2405–2415
5. Wu P, Sun P, Nie KX, Zhu YB, Shi MY, Xiao CG, Liu H, Liu QY, Zhao TY, Chen XG, Zhou HN, Wang PH, Cheng G#. (2019) A gut commensal bacterium promotes mosquito permissiveness to arboviruses. **Cell Host & Microbe,** 25, 1-12.
6. Liu Y, Liu JY, Du SY, Shan C, Nie KX, Zhang RD, Li XF, Zhang RL, Wang T, Qin CF, Wang PH, Shi PY#, Cheng G#. (2017) Evolutionary enhancement of Zika virus infectivity in *Aedes aegypti* mosquitoes. **Nature.** 545: 482–486.
7. Cheng G, Cox J, Wang PH, Krishnan MN, Dai JF, Qian F, Anderson JF, Fikrig E. (2010) A C-Type Lectin collaborates with a CD45 phosphatase homolog to facilitate West Nile Virus infection of mosquitoes. **Cell.** 142, 714–725.

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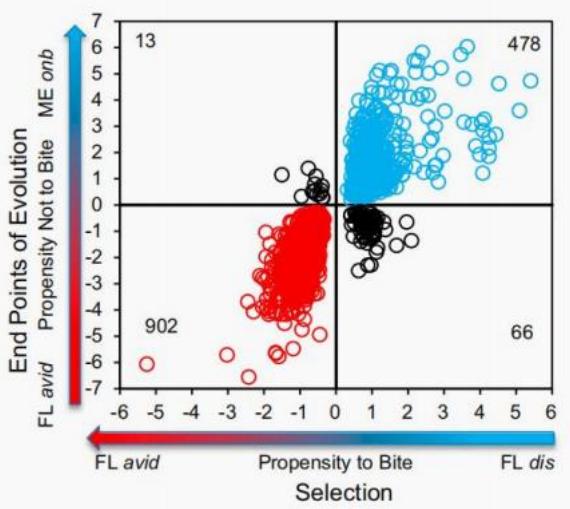
Evolutionary transition from blood feeding to obligate nonbiting in a mosquito



紫色猪笼草蚊

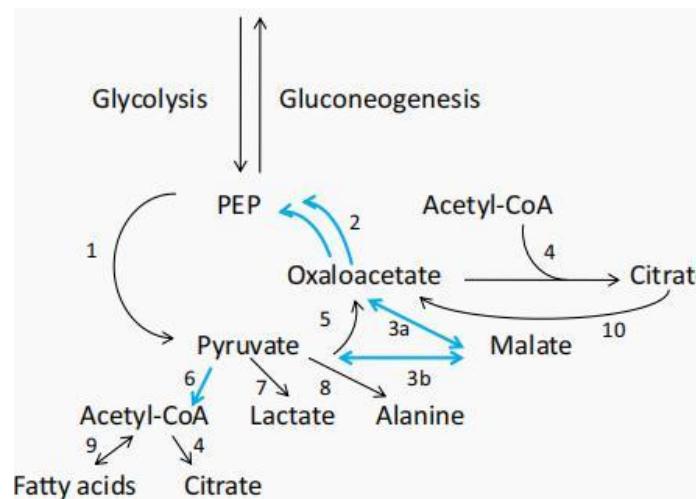
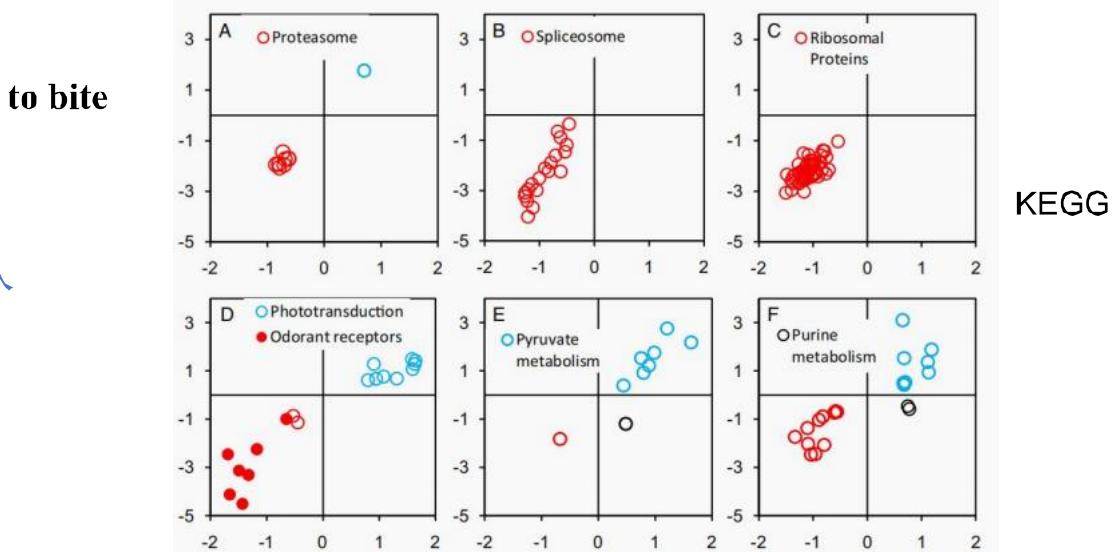
Florida population: a low propensity to bite
avid biters (FLavid) 强制
disinterested nonbiters (FLdis)
Maine population:
obligate nonbiters (MEonb) 专性非咬人

differential gene expression (DGE)



anautogenous: require blood to develop eggs

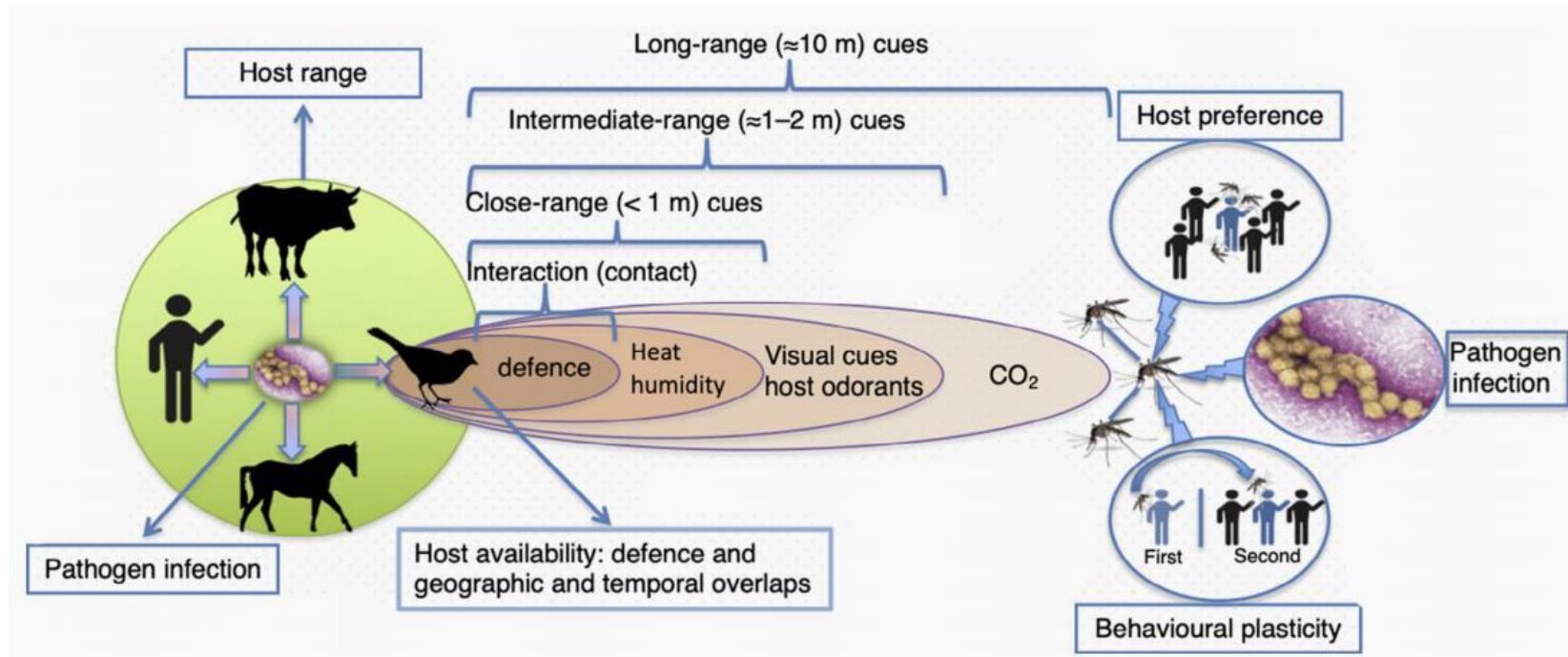
autogenous: producing one or more clutches of eggs without blood



nonbiting :
a greatly reduced metabolic investment,
greater reliance on visual rather than olfactory sensory input

Summary:

Mosquito host utilization is a highly complex phenomenon influenced by both intrinsic and extrinsic factors



Host preferences: limit what type of hosts mosquitoes feed on

Behavioural plasticity: enable shifting from a preferred host to an available one

Host-seeking cues and host availability: determine host–vector interactions

Pathogen infection: alter host and vector phenotypic traits

Yan J, Gangoso L, et al., *Biol Rev Camb Philos Soc*, 2021

Take home message

- Sugar feeding is also important for mosquitoes, which search for nectar through visual, smell, taste and other clues, but the neuromolecular mechanism behind it has been less studied
- Some ways to study mosquito feeding behavior, such as Y-tube、three-cage olfactometers and two-port olfactometers
- Mosquitoes use multiple senses, including smell, sight, heat reception, humidity sensing, and taste, to track their hosts
- The mosquito's preference for the host can be changed by learning, being infected by viruses, etc.

Part II: The internal mechanisms govern feeding

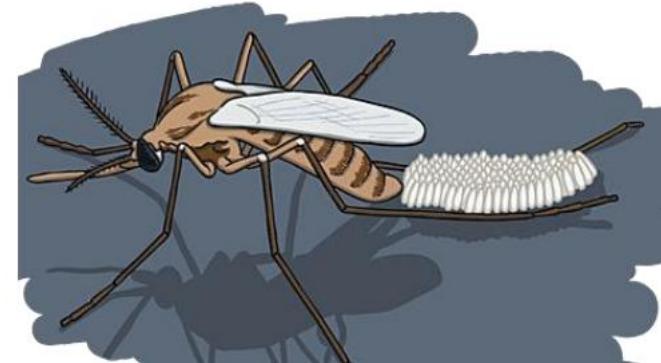
JSH

Q1: What drives mosquitoes to start **blood** feeding?

Q2: How can mosquitoes keep **blood** feeding for a while?

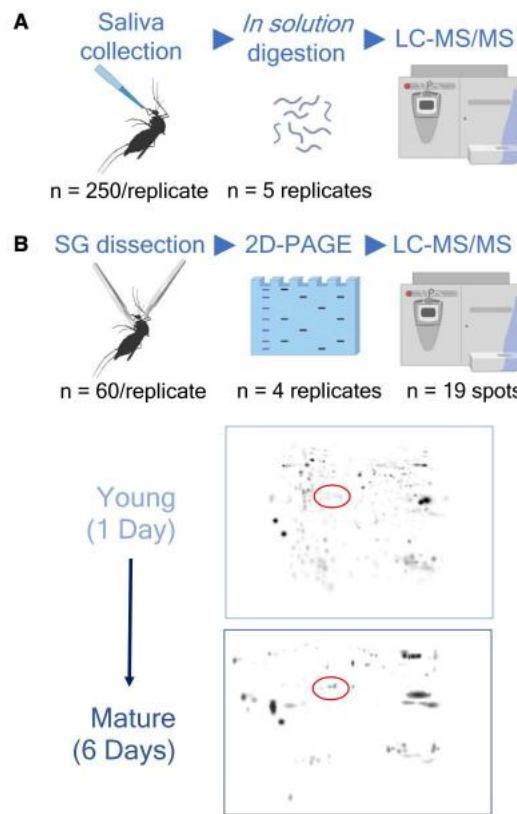
Q3: How do mosquitoes stop **blood-feeding**?

Risks always coexist with opportunities

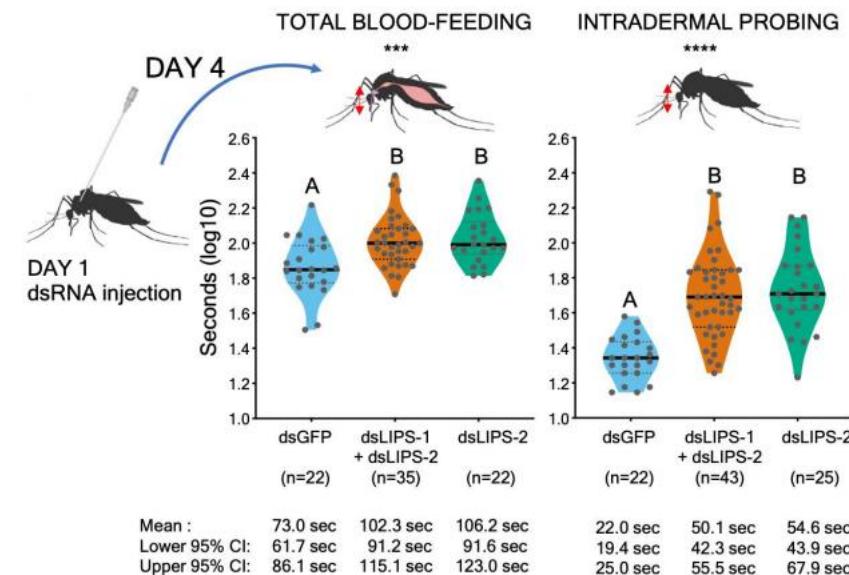
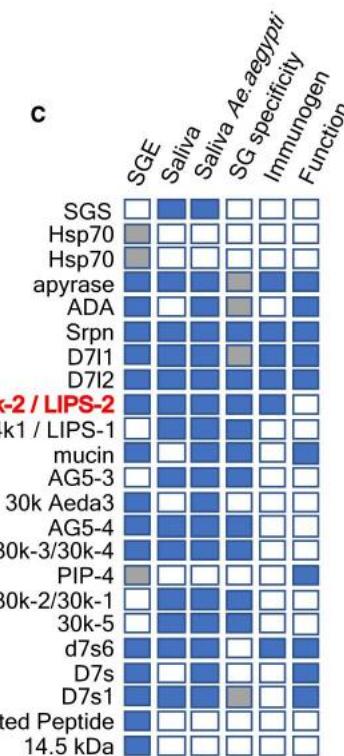




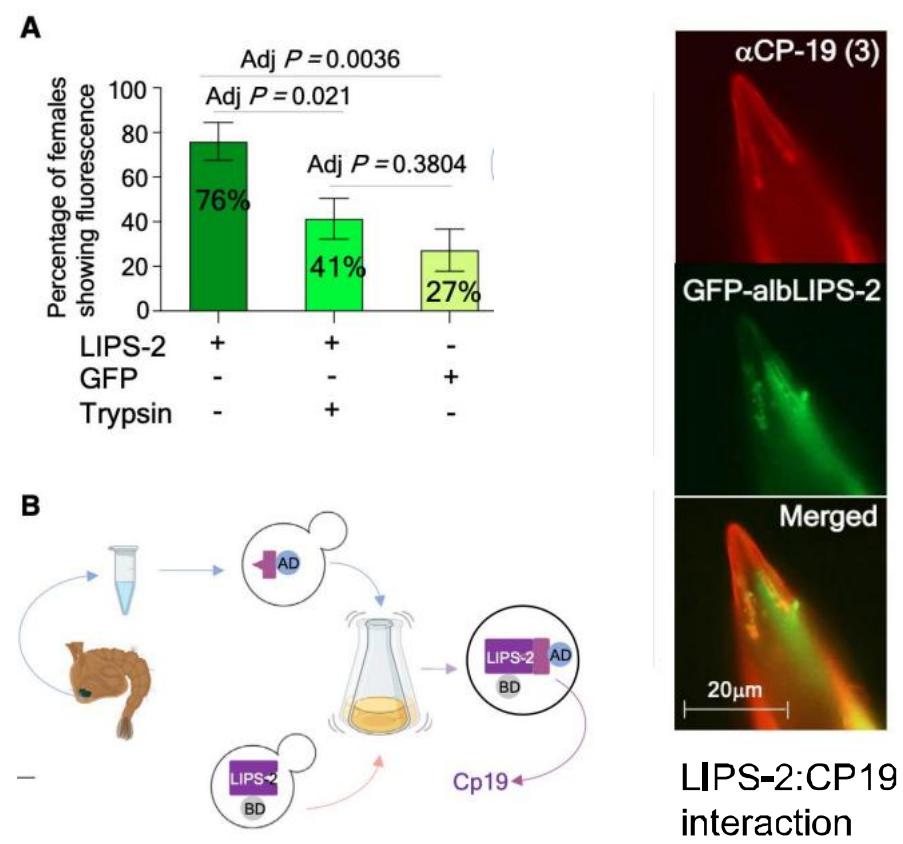
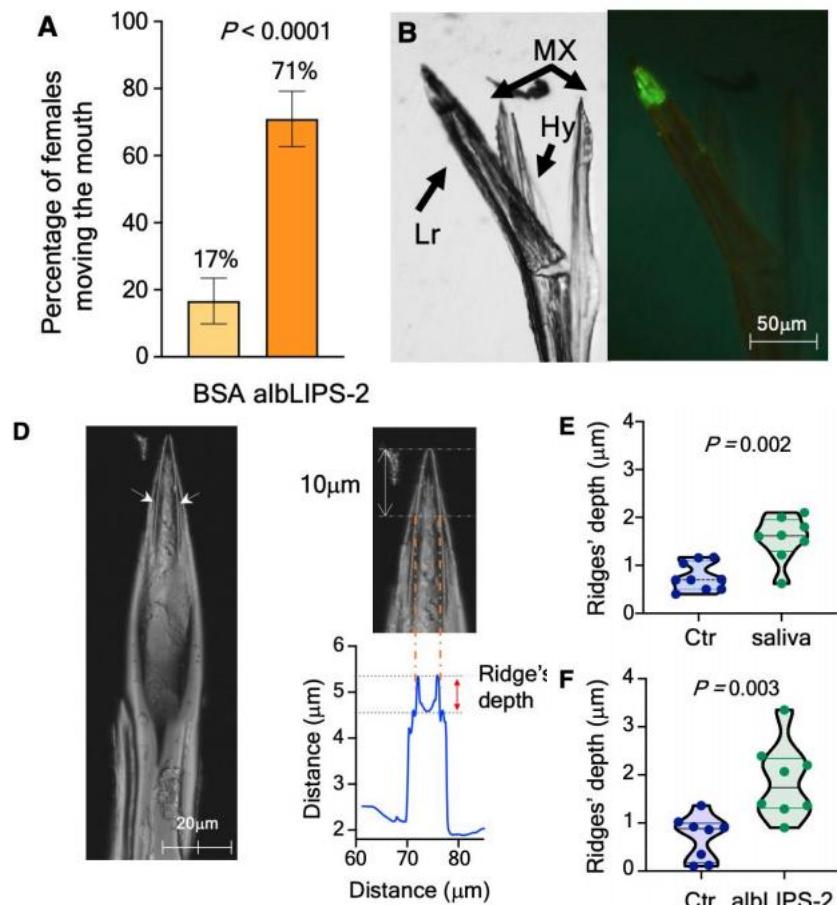
Blood-feeding initiation



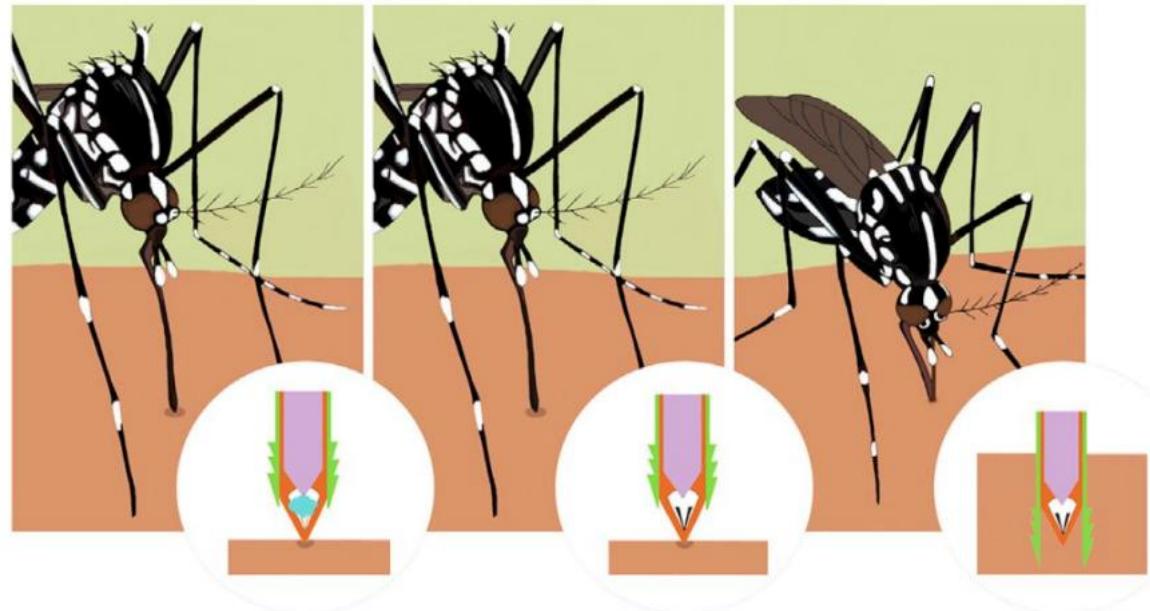
The presence of LIPS proteins in *Aedes saliva* is critical for biting



LIPS-2 binding Cp19 causes a morphological change of the structures of the labrum

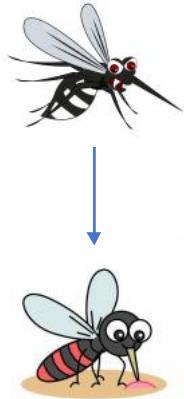


Initiation of Blood-feeding

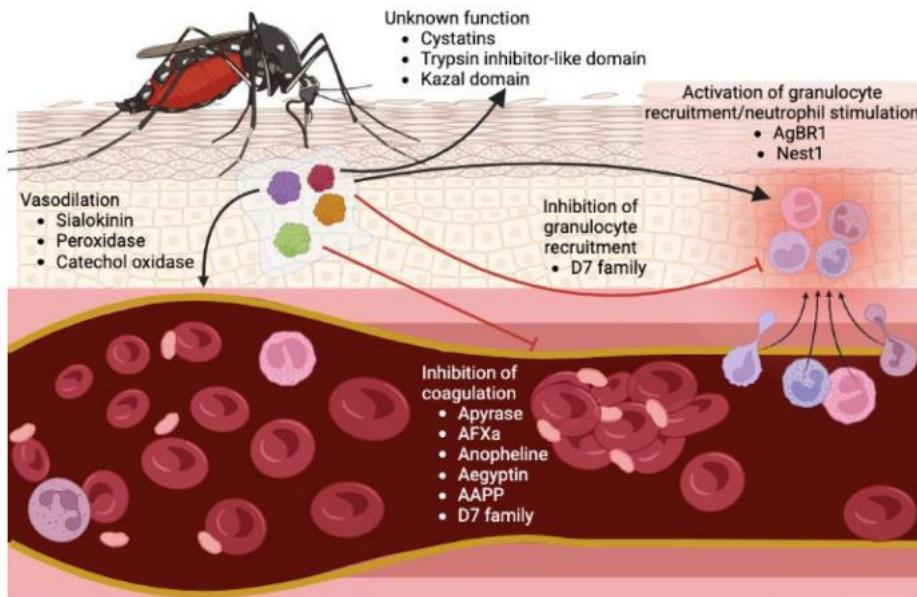


Arnoldi I, et al. Current Biology. 2022

Salivary proteins facilitate mosquito blood feeding



Blood-feeding



Marín-López, A, et al. *Pathogens*. 2023

Host hemostasis:

- platelet aggregation
- vascular contraction
- blood clotting

Salivary proteins:

- D7 protein family
- sialokinin
- aegyptin
- adenine deaminase
- ...

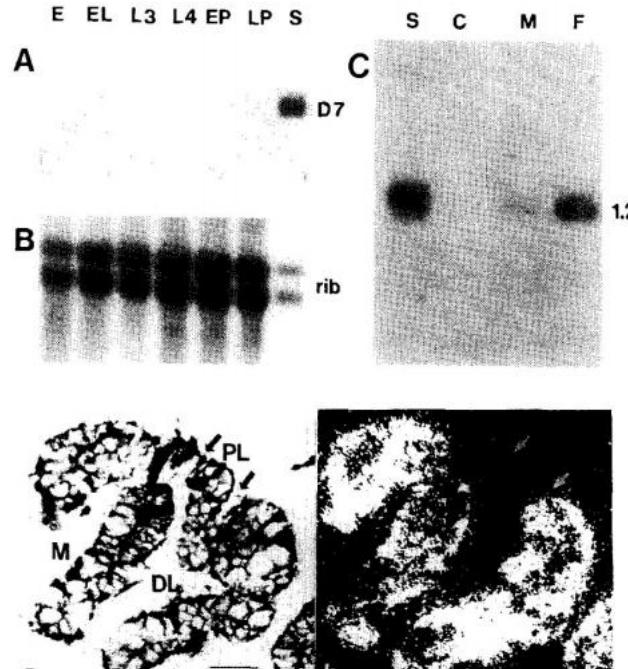
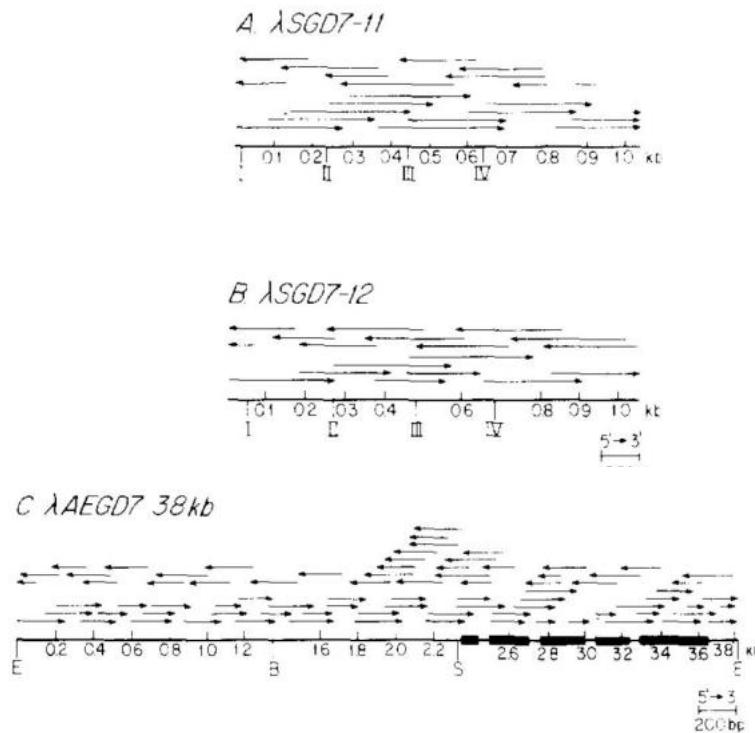
Isolation and characterization of the gene expressing the major salivary gland protein of the female mosquito, *Aedes aegypti*

Anthony A. James, Karen Blackmer^{*1}, Osvaldo Marinotti^{*2}, Corine R. Ghosn, and Jeffrey V. Racioppi^{*3}

Department of Molecular Biology and Biochemistry, University of California, Irvine, CA, U.S.A.

(Received 13 July 1990; accepted 27 August 1990)

"sex- and stage-specificity"



The feature of D7 Proteins:

- 1) widespread in hematophagous Diptera
(exclusively expressed in female salivary glands)
- 2) different length and amino acid sequence
(in different species and within the same species)
- 3) binding small hydrophobic molecules
(host responses : inflammation and hemostasis)

Ae. aegypti D7L1
(AAEL006424)

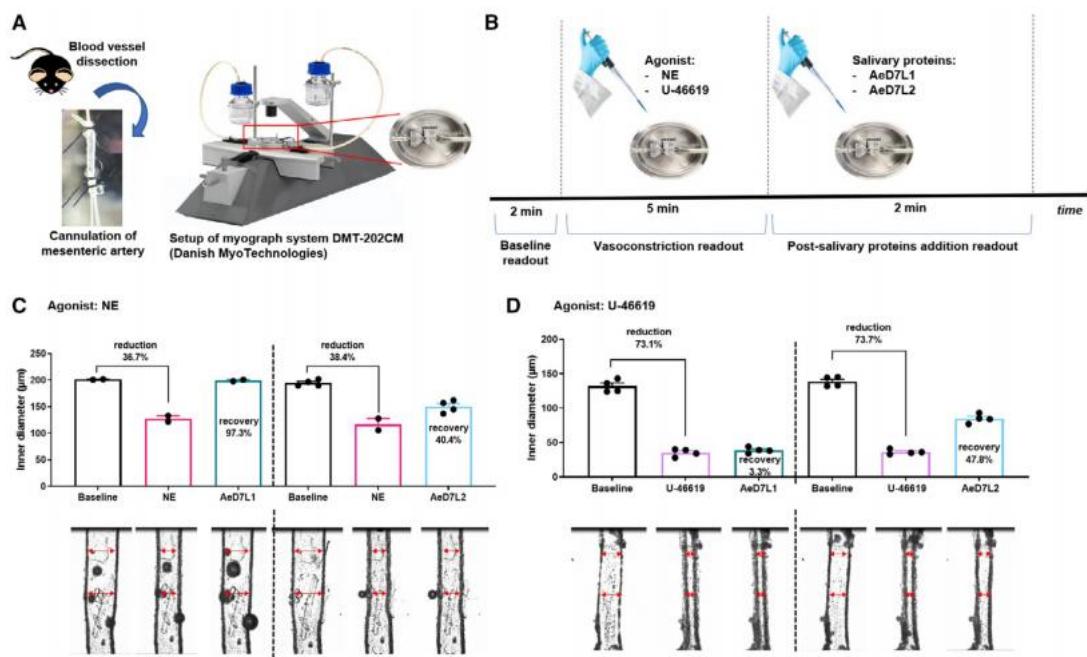
Ae. aegypti D7L2
(AAEL006417)

	**	3DXL
Serotonin (0.39)		
Norepinephrine (0.119)	3DYE	
Histamine (140)		
Epinephrine (102)		
LTC4 (57.4)		
LTD4 (54.3)		
LTE4 (60.2)	3DZT	
LTB4 (140)		
No binding to U46619	TxA2	
Serotonin (1.68)		
Norepinephrine (110)		
Histamine (1130)		
LTC4 (5270)		
LTD4 (597)		
LTE4 (1930)		
LTB4 (769)		

Alvarenga, P.H. and Andersen, J.F. Biology. 2023

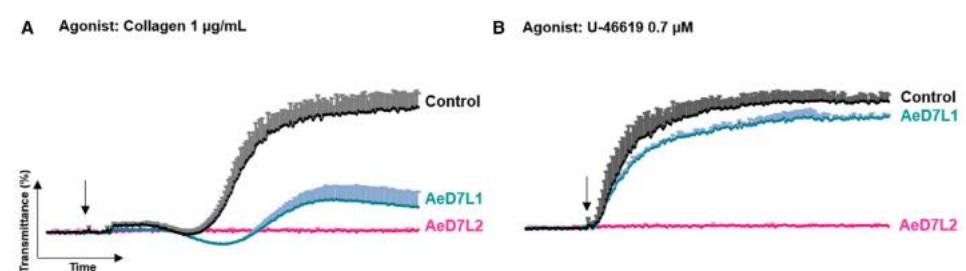
AeD7L1 and AeD7L2 interfere with vasoreactivity and platelet aggregation without compromising blood coagulation

vascular contraction

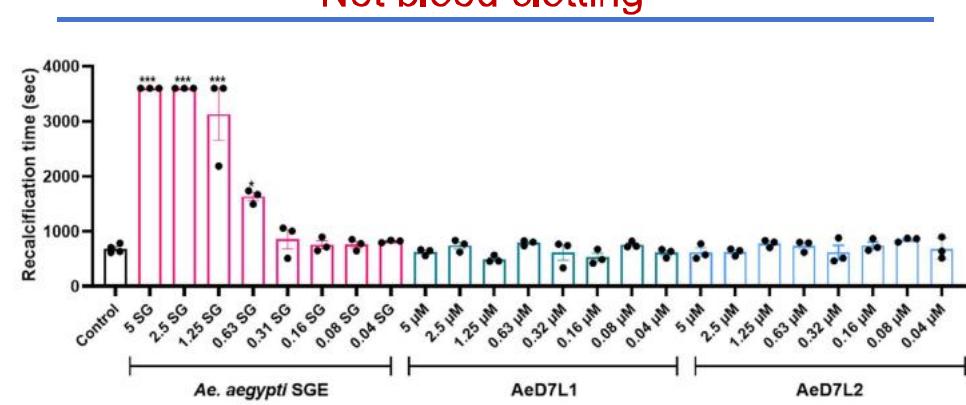


I. Martin-Martin et al. The FEBS Journal, 2021

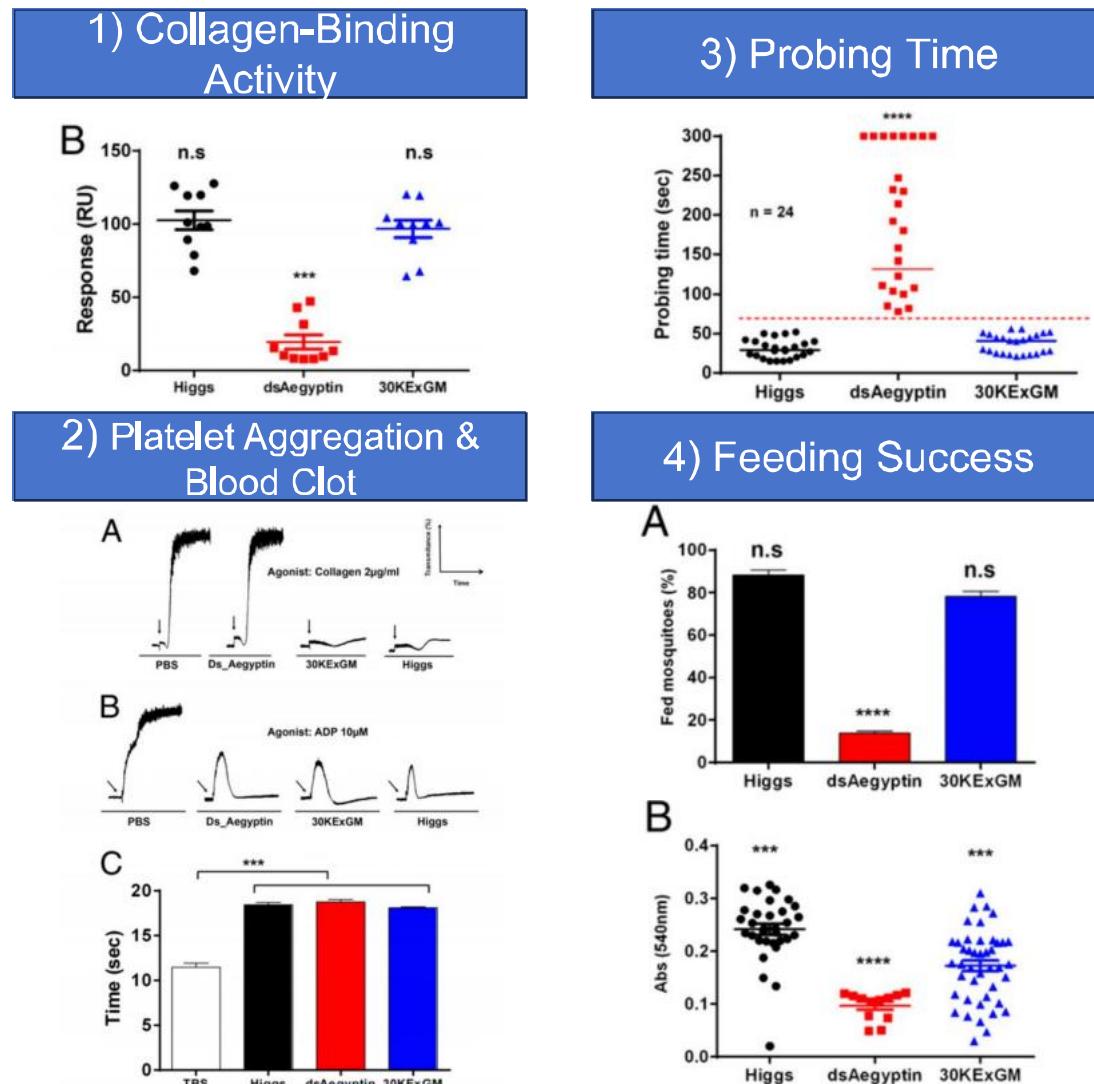
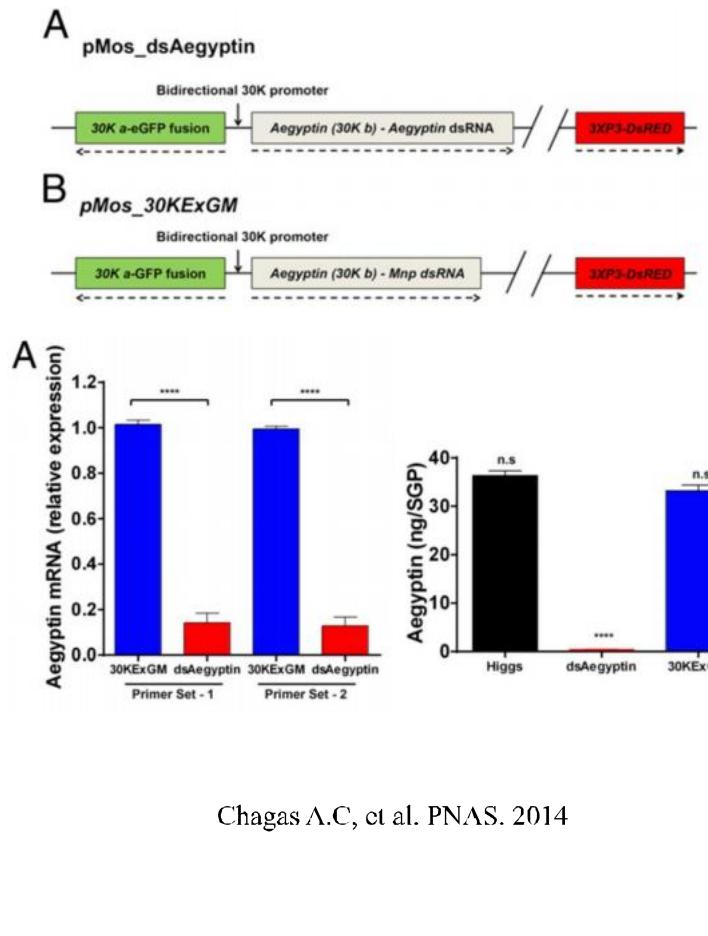
platelet aggregation



Not blood clotting



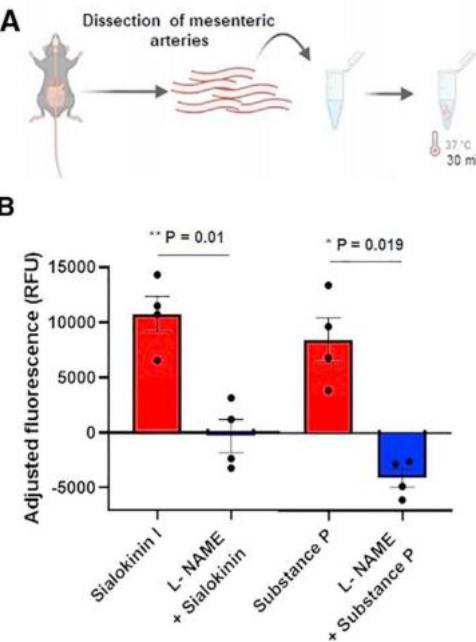
Aegyptin regulates probing time and blood feeding success via binding collagen



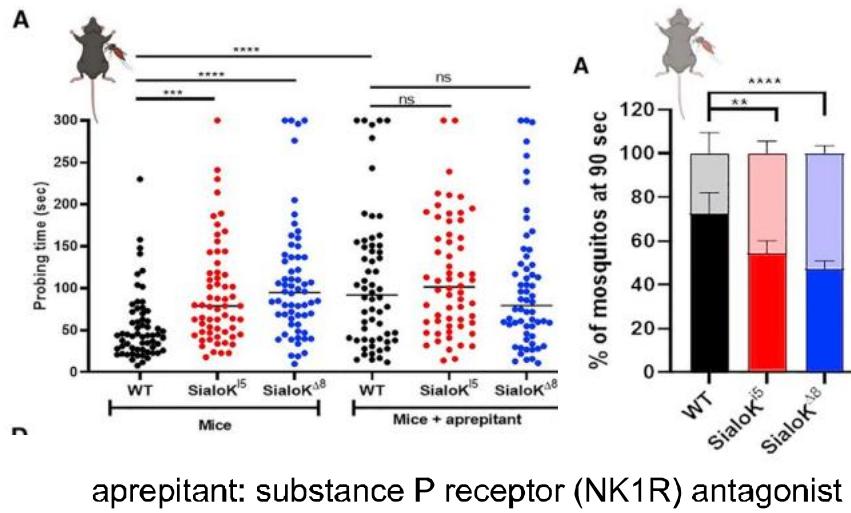
Chagas A.C, et al. PNAS. 2014

Sialokinin facilitates mosquito blood feeding

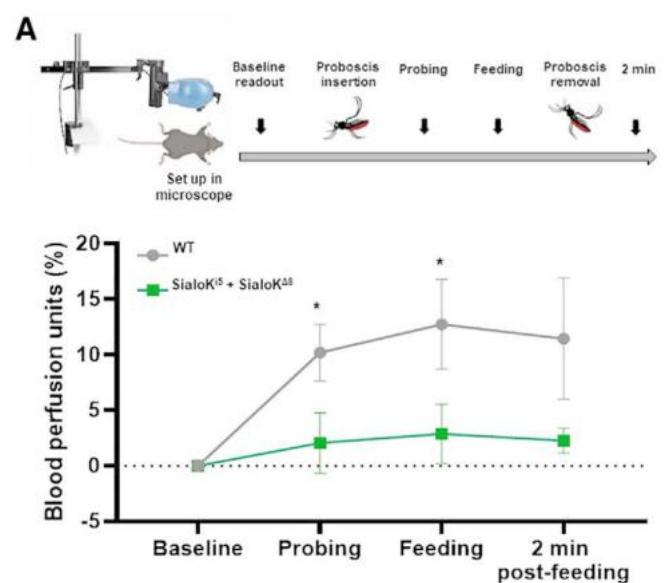
Induce nitric oxide release



Interferes with host hemostasis via NK1R signaling



Promotes blood perfusion at the bite site

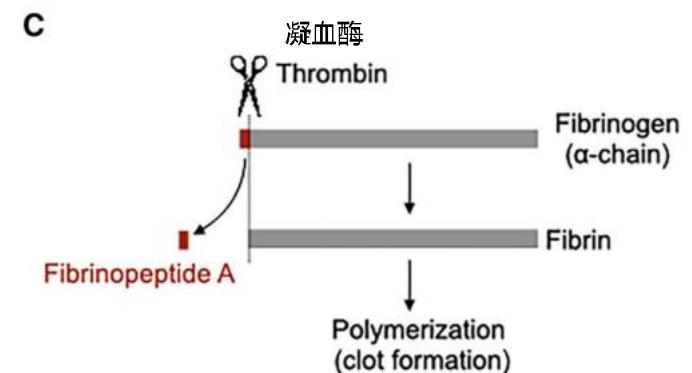
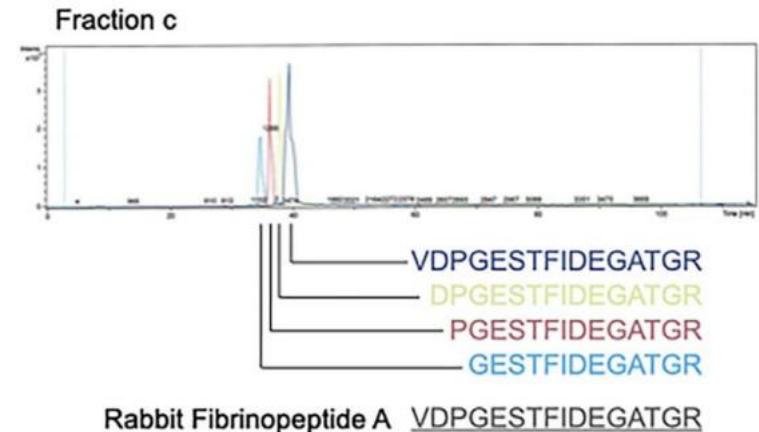
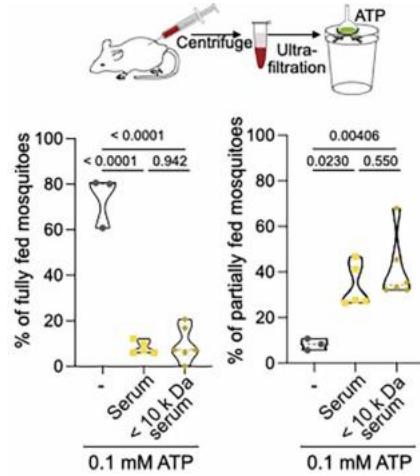
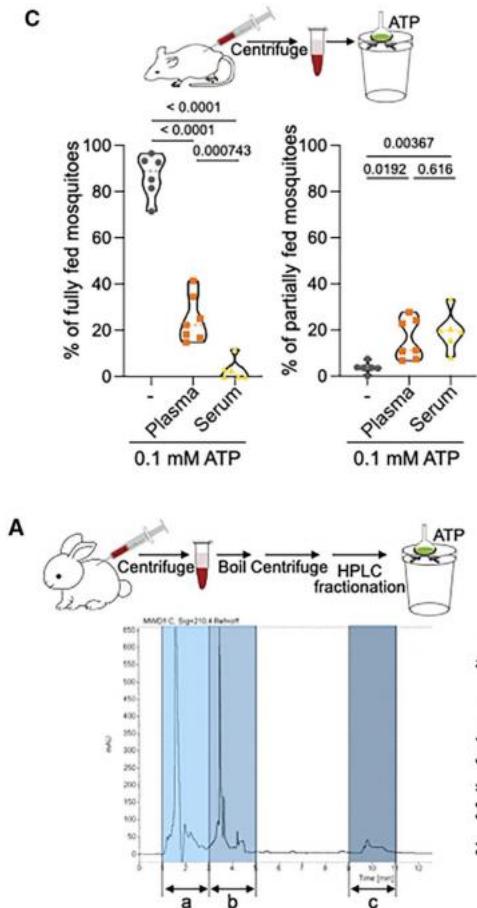


Ensures short probing time and successful blood feeding!

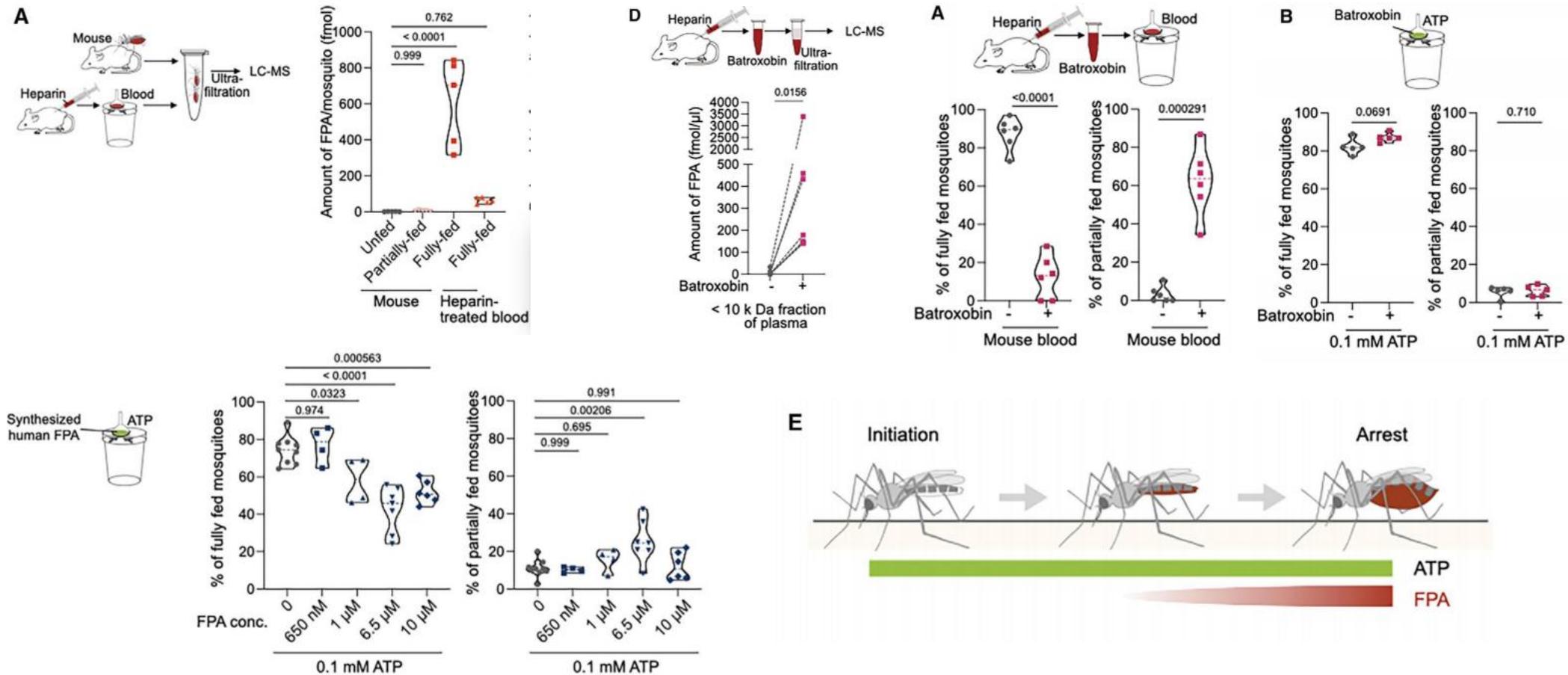
FPA from blood serum as a potential factor inhibiting blood feeding



Blood-feeding
termination



FPA arrests the process of mosquito blood feeding



Abdominal distension may participate in blood-feeding arrest

REGULATION OF BLOOD MEAL SIZE IN THE MOSQUITO*

ROBERT W. GWADZ

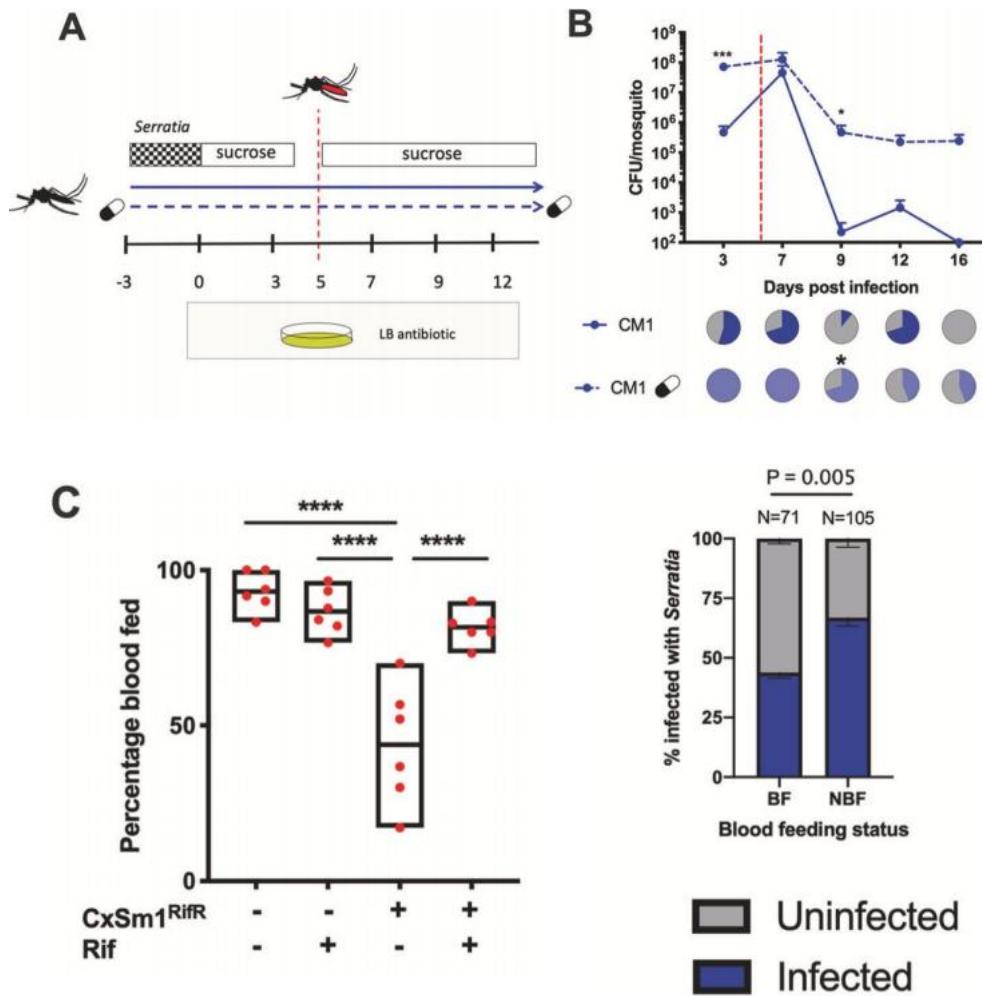
Department of Biology, University of Notre Dame,
Notre Dame, Indiana 46556

(Received 14 June 1969)

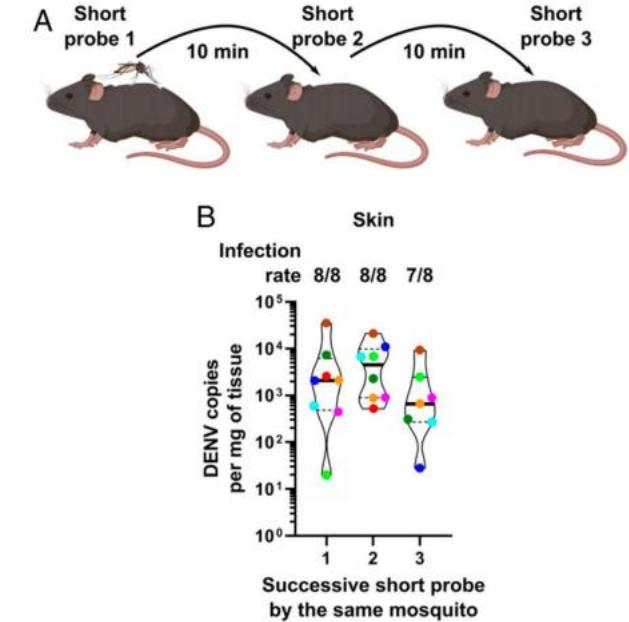
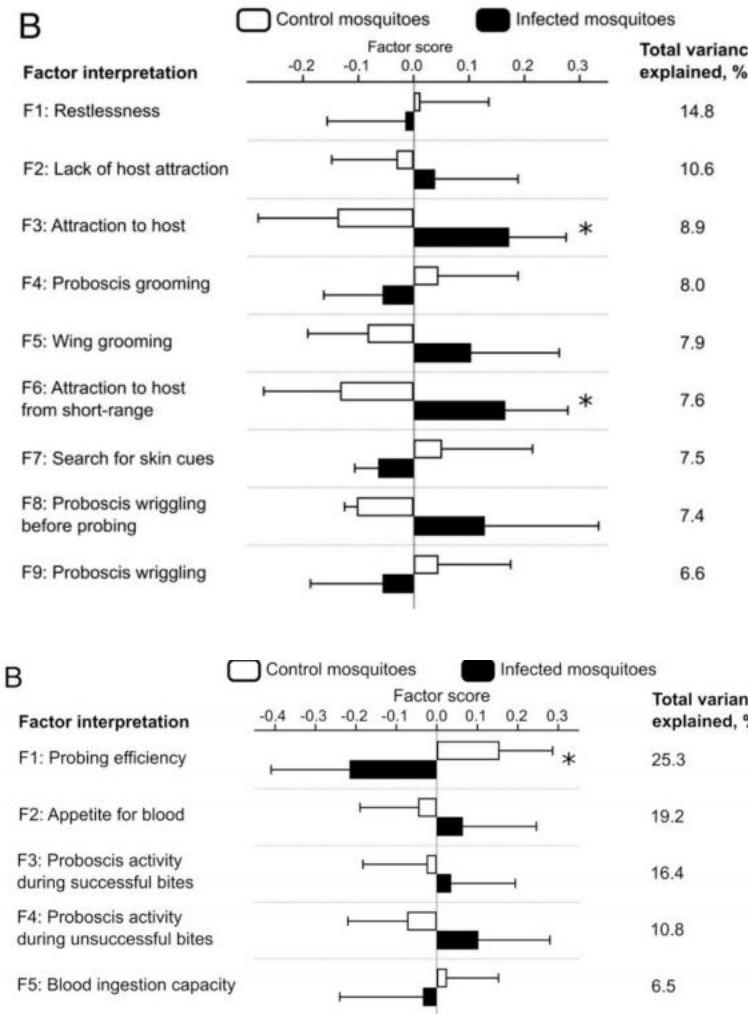
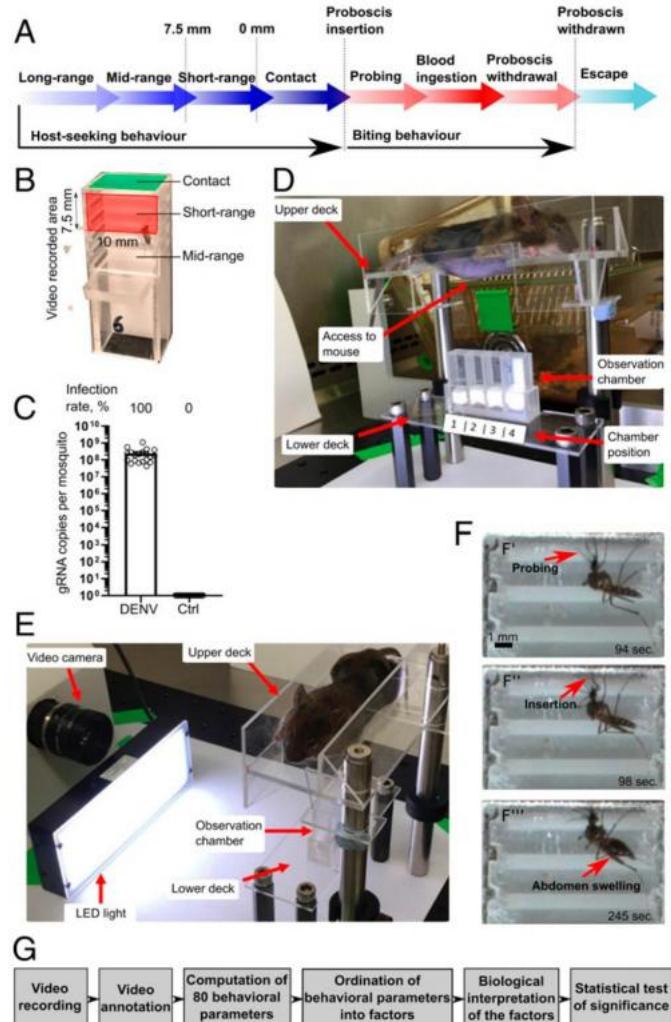
Treatment: Ventral nerve cord cut anterior to abdominal ganglion	No. females/ treatment	Mean for treated females (\pm 95% confidence limits)		
		Feeding time (min)	Blood meal weight (mg)	No. of eggs developed
No. 2	76	5.31 \pm 0.24	11.99 \pm 0.51	103.28 \pm 4.31*
No. 3	42	3.21 \pm 0.32	7.63 \pm 0.70	97.17 \pm 4.01
No. 4	36	2.69 \pm 0.22	5.14 \pm 0.40	92.32 \pm 2.81
No. 5	40	2.19 \pm 0.16	4.23 \pm 0.46	88.88 \pm 4.25
No. 6	36	1.98 \pm 0.18	3.61 \pm 0.21	87.14 \pm 4.39
Sham-operated†	62	1.76 \pm 0.11	2.93 \pm 0.18	85.78 \pm 3.03
No treatment‡	64	1.79 \pm 0.10	2.85 \pm 0.26	86.07 \pm 3.30



Microbial interactions in the mosquito gut determine blood-feeding behavior

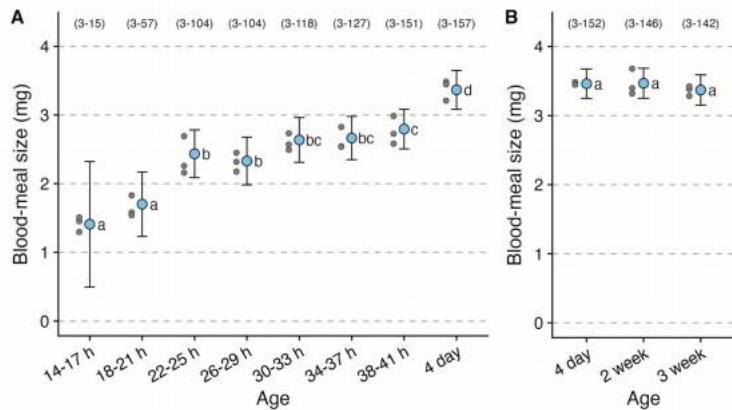


DENV Infection increases host attraction and frequency of infectious probes

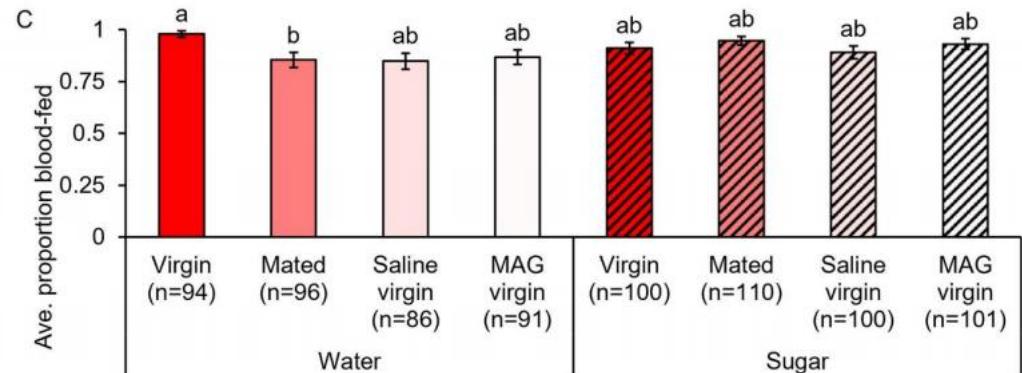


WeiXiang, et al. PNAS. 2022

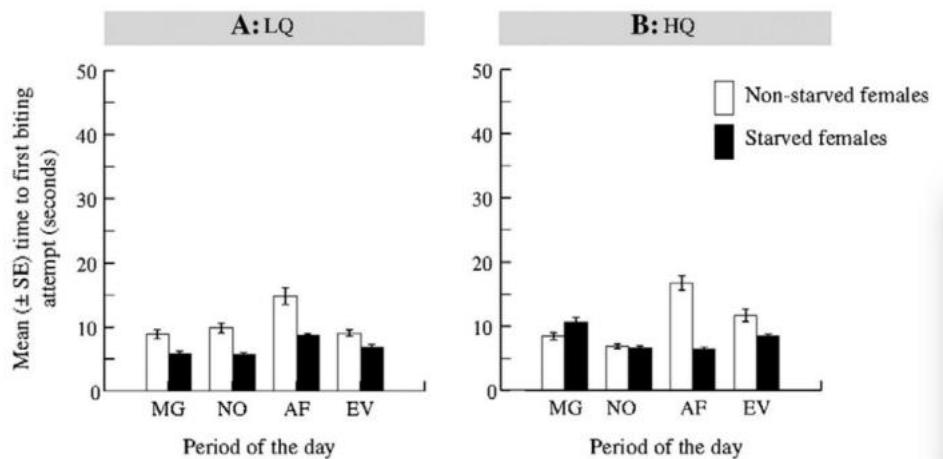
Physiological relevance in blood feeding



Ramírez-Sánchez L.F. et al. Journal of Insect Physiology. 2023



League G.P., et al. PLOS NEGLECTED TROPICAL DISEASES. 2021



Dieng H., et al. Journal of Asia-Pacific Entomology. 2015

Summary:

- The LIPS-2 in the saliva binding Cp19 causes a morphology change of the structures of the labrum.
- Saliva contains many bioactive molecules that promote blood feeding by inhibiting hemostasis.
- There are negative controls arresting blood-feeding. eg. FPA, stretch receptors.
- Bacterial flora and disease pathogens change probing efficiency.
- Physiological state affects blood feeding.

Mechanisms of external factors affecting mosquito feeding

姜思梅

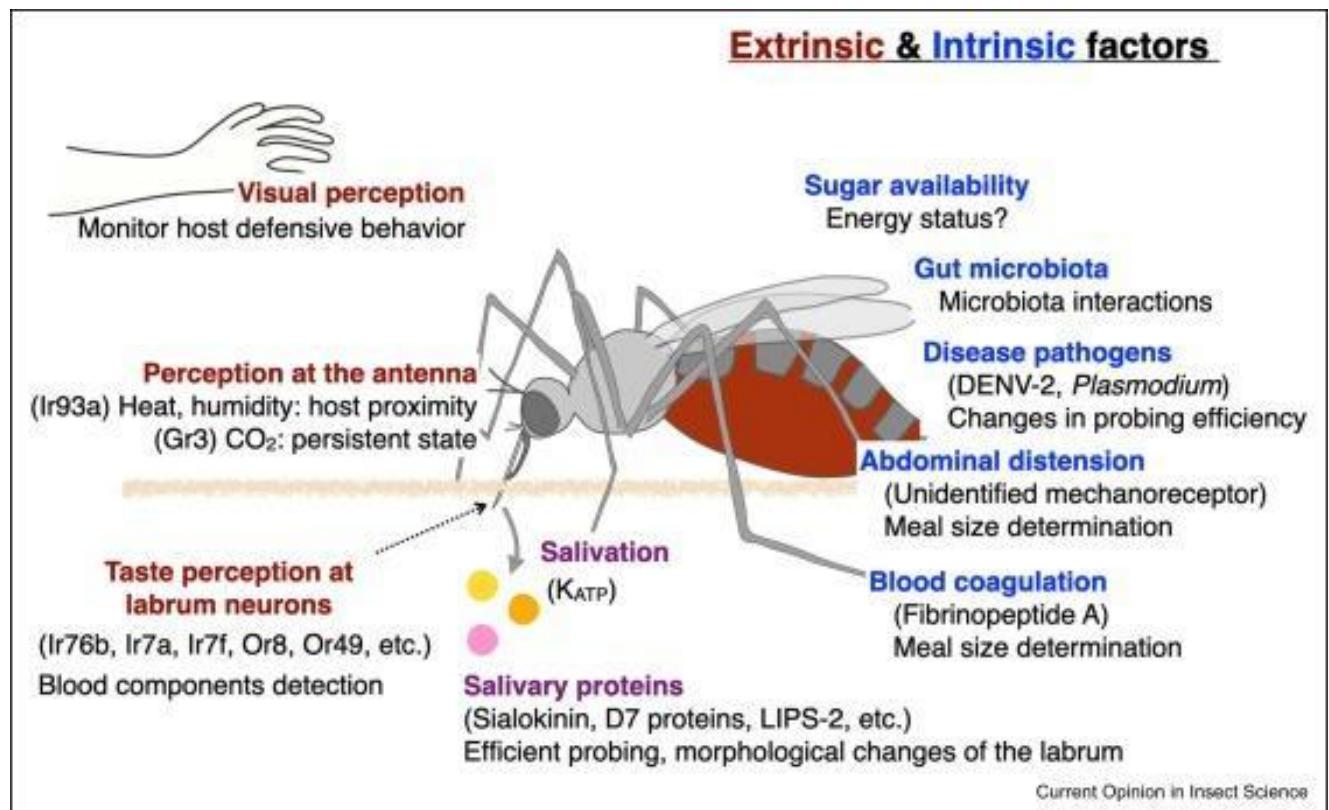
2024.9.26

Contents

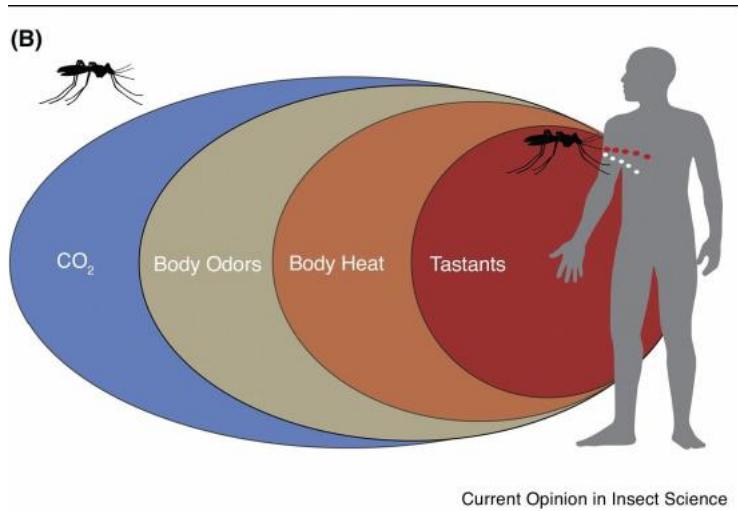
1. Which external factors affect mosquito feeding?
2. How do these factors affect mosquito feeding?
3. The most famous mosquito repellent - DEET

Which external factors affect mosquito feeding?

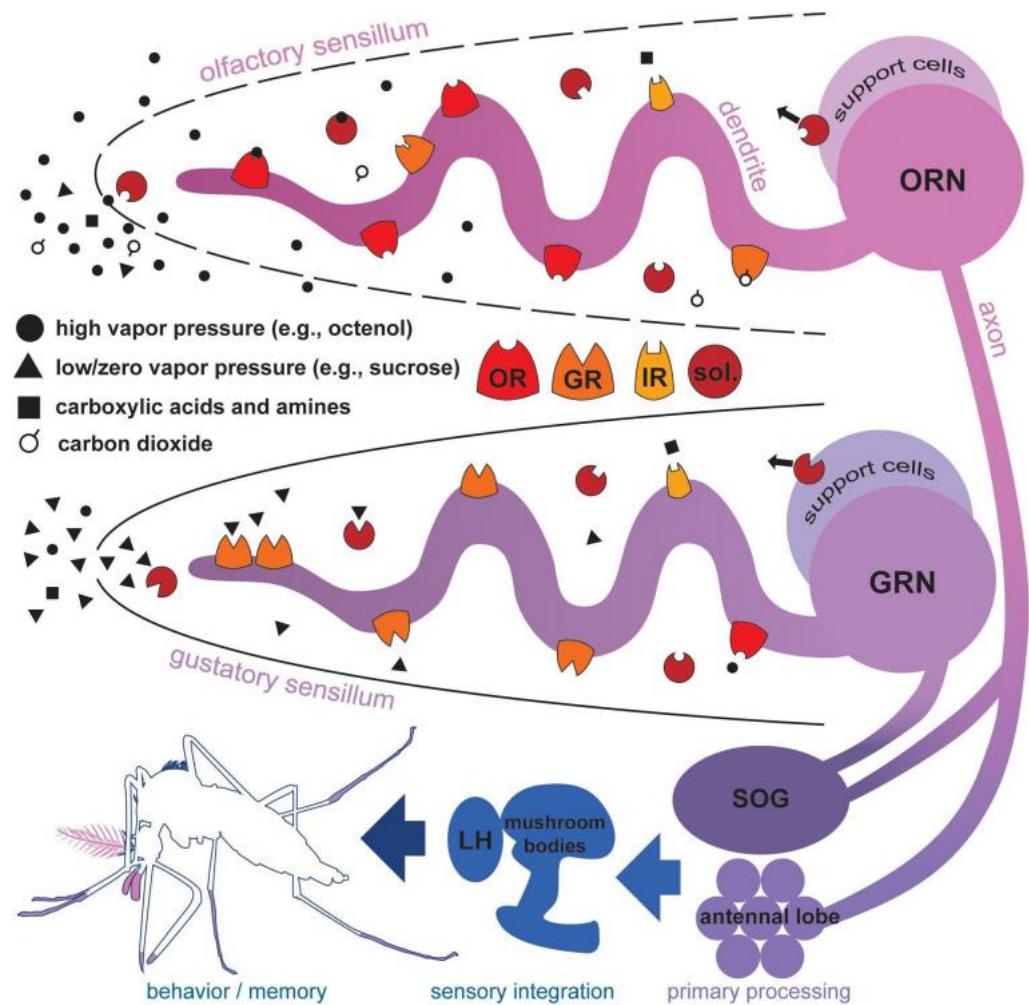
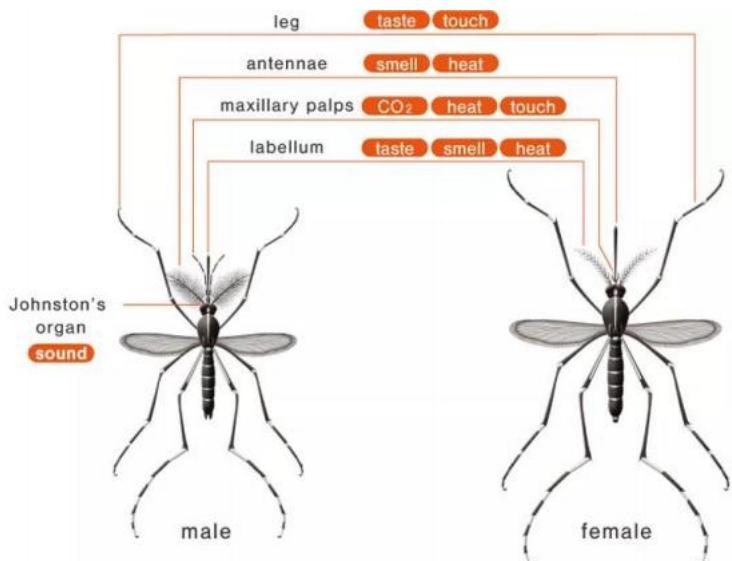
- Vision: host defensive behavior
- Olfaction: host's skin odour, CO₂
- Gustation: the taste and the quality of host blood
- Environmental factors: temperature, humidity, light



How do these factors affect mosquito feeding?



Raji JL, et al. *Curr Opin Insect Sci.* 2017

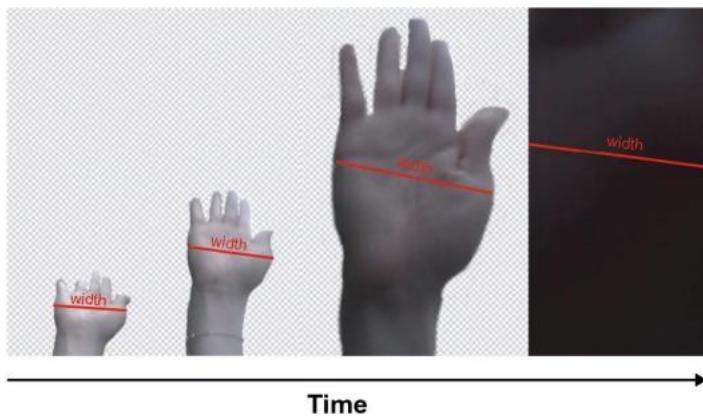


Sparks JT, et al. *Front Physiol.* 2018

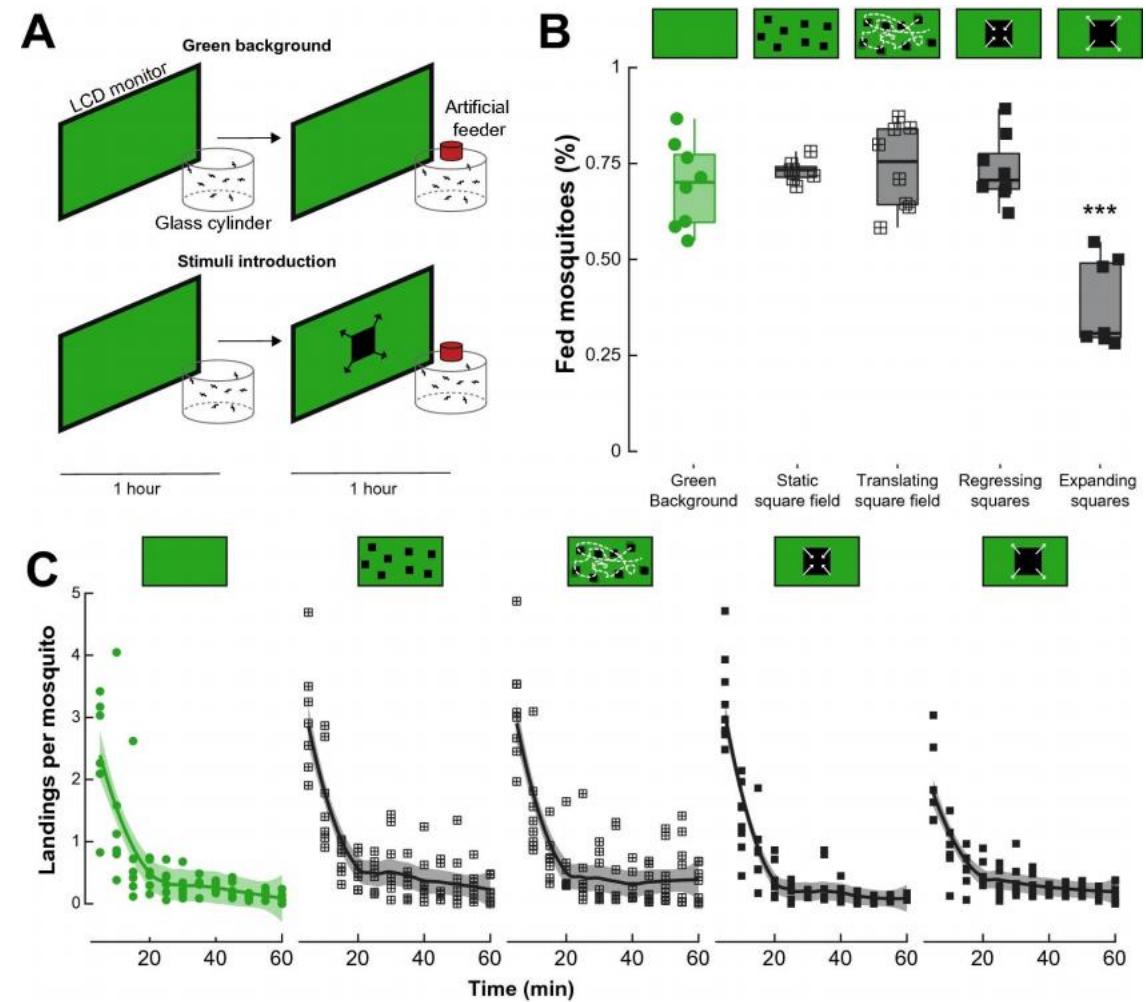
• Vision

**Visual motion reduces feeding, but not landing in
*Aedes aegypti***

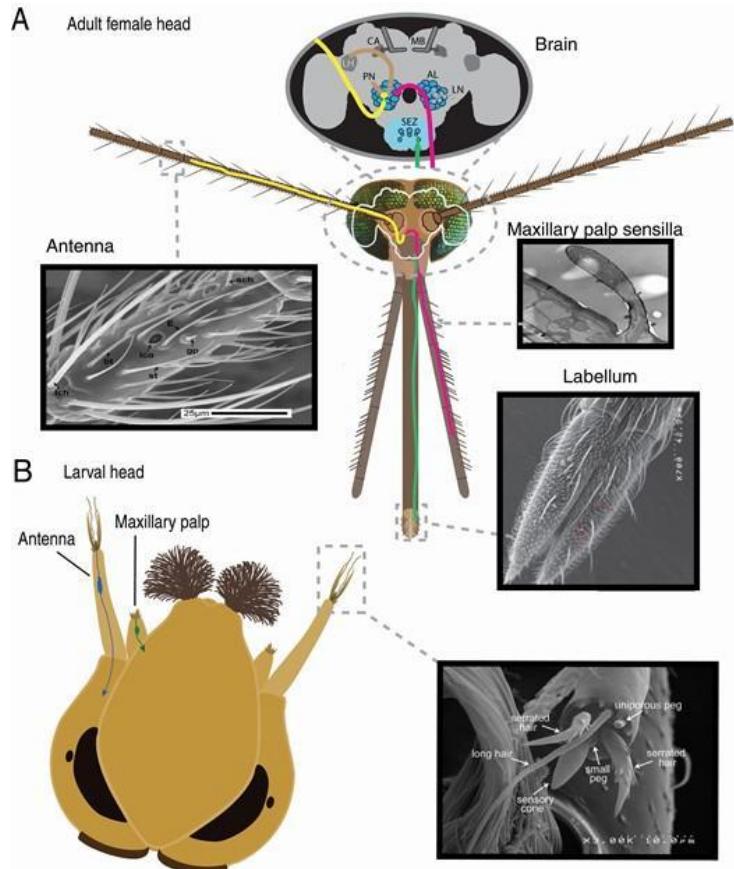
拍手 {
 空气流动（机械成分）
 视野中物体的快速扩张
 （视觉成分）



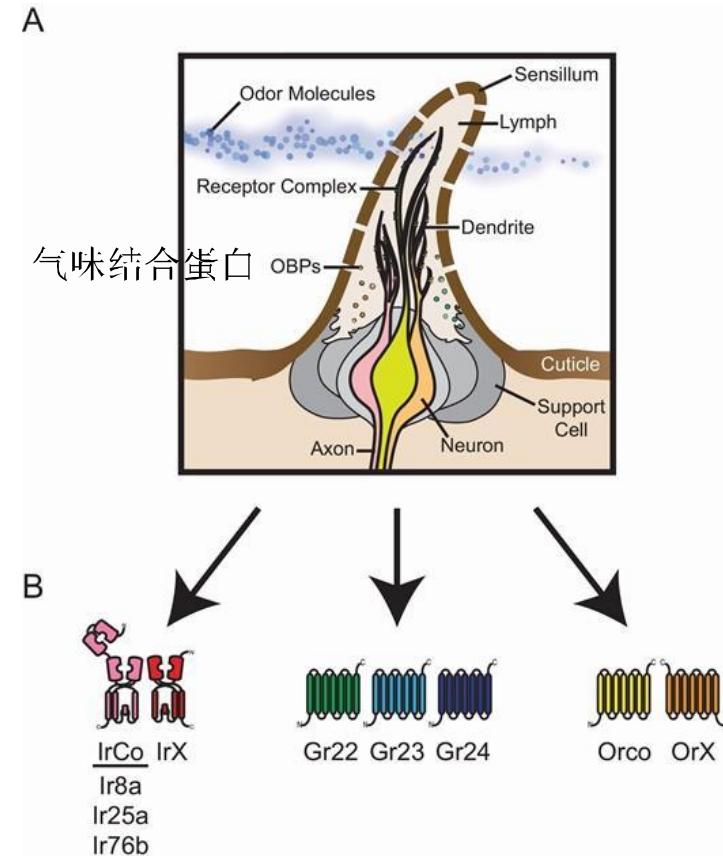
Wynne NE, et al. Sci Rep. 2022



• Olfaction



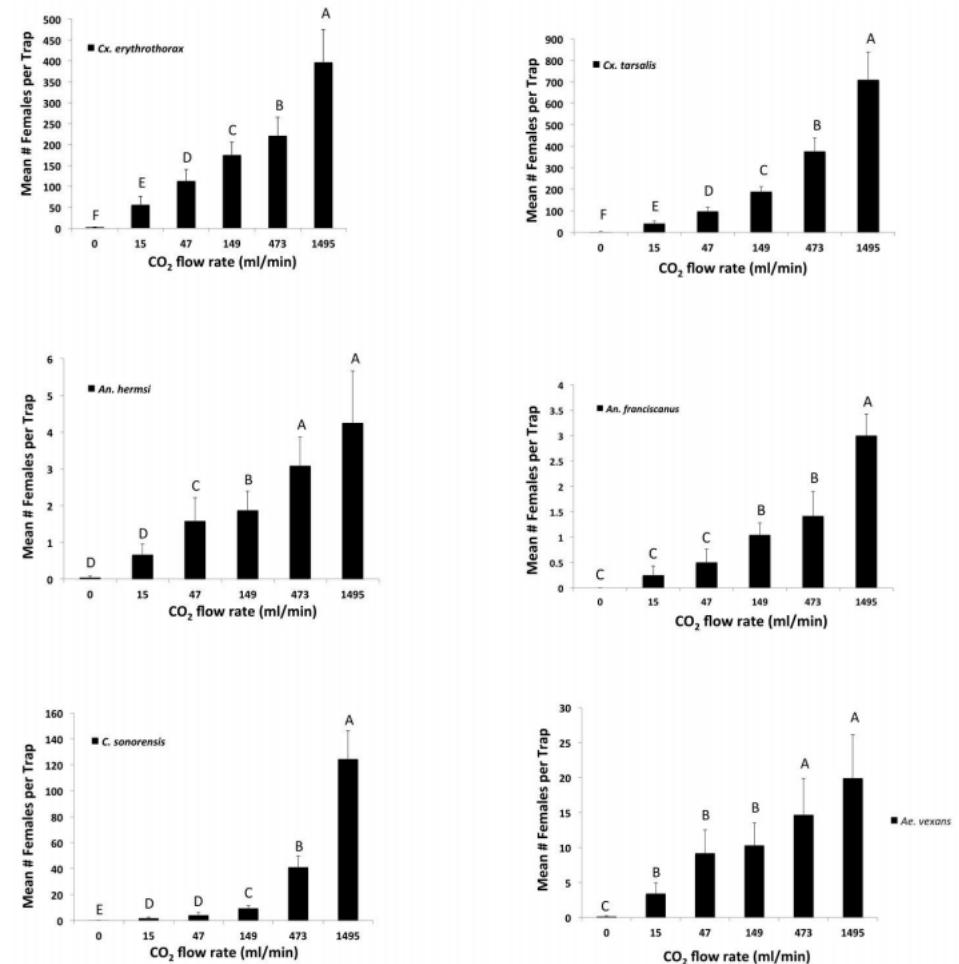
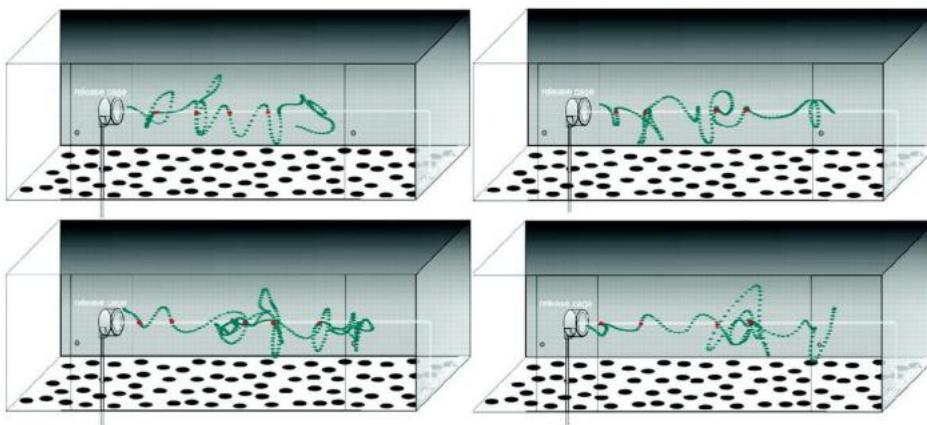
按蚊嗅觉系统示意图



按蚊的感觉器和嗅觉受体的解剖结构

Konopka Joanna K, et al. *Chem Senses*. 2021

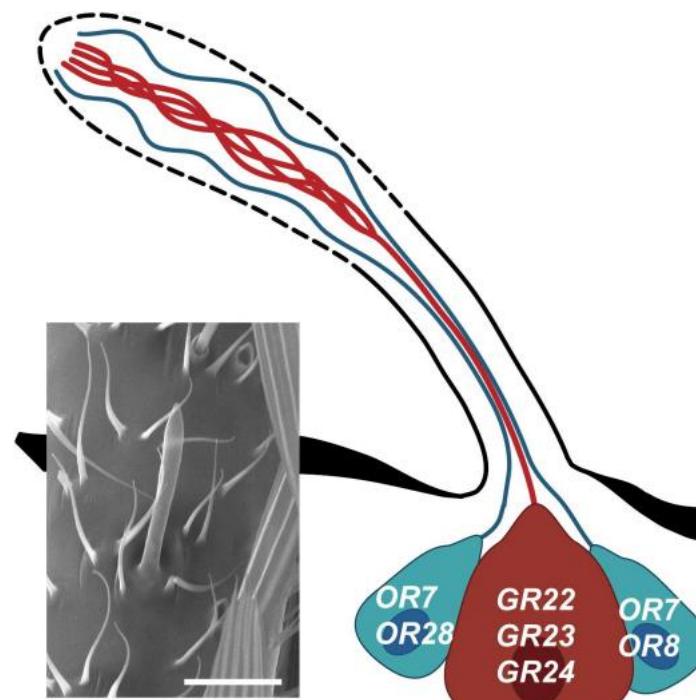
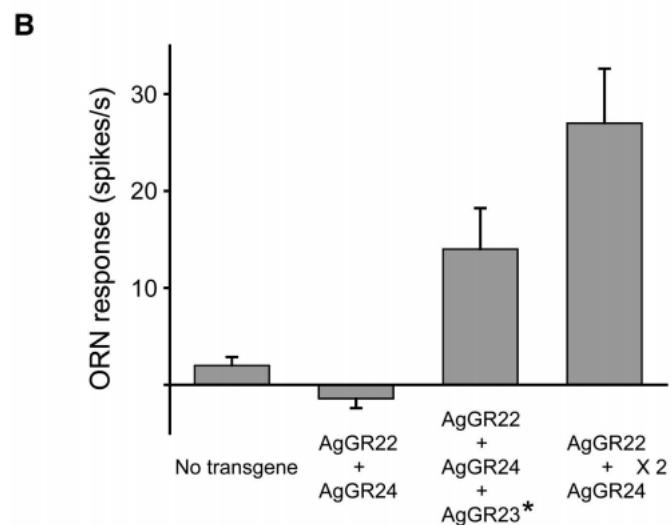
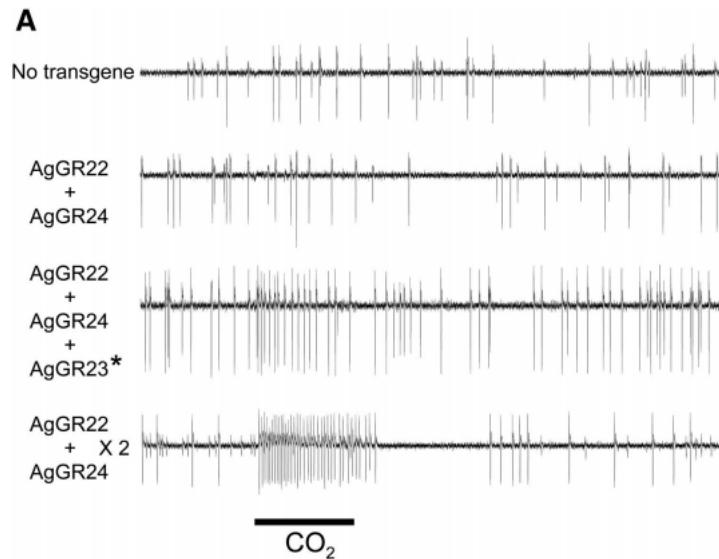
CO_2 is highly attractive to mosquitoes and causes behavioral changes



Dekker T, et al. *J Exp Biol.* 2011

McPhatter L, et al. *J Vector Ecol.* 2017

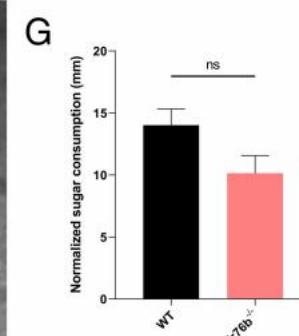
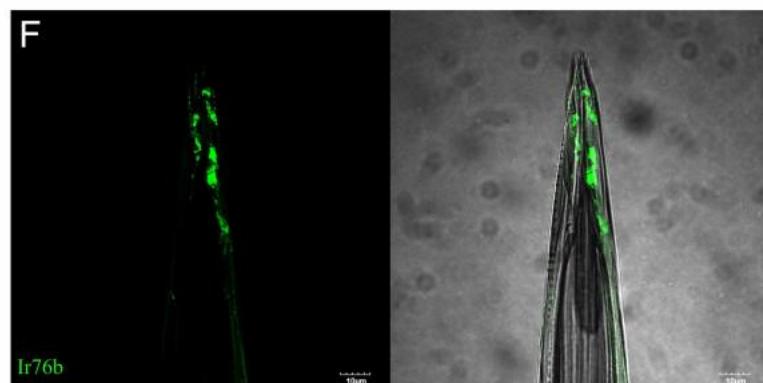
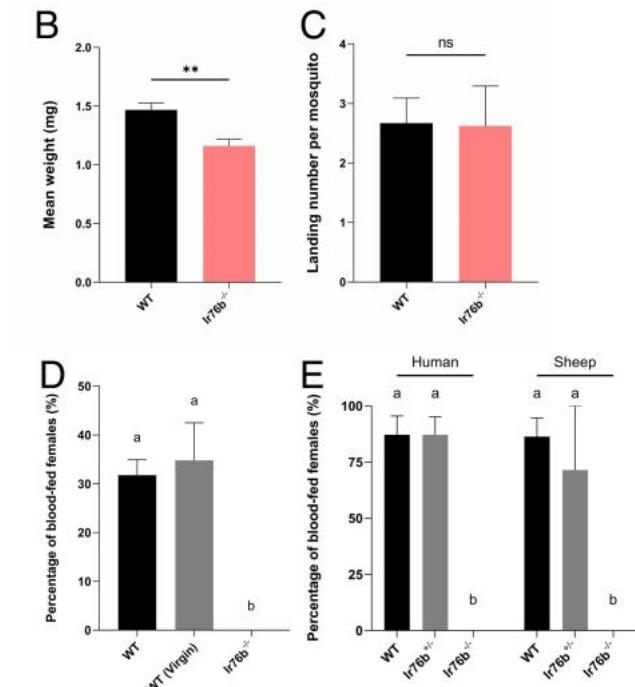
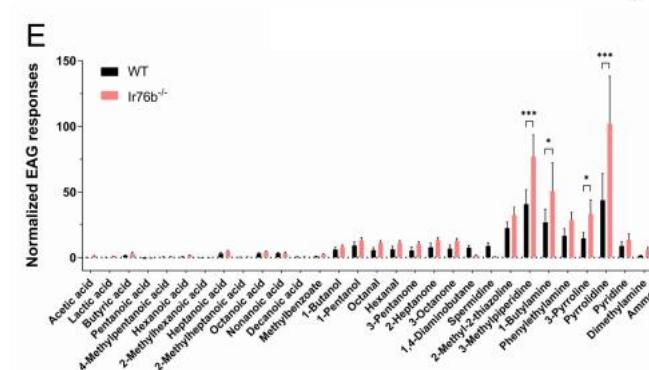
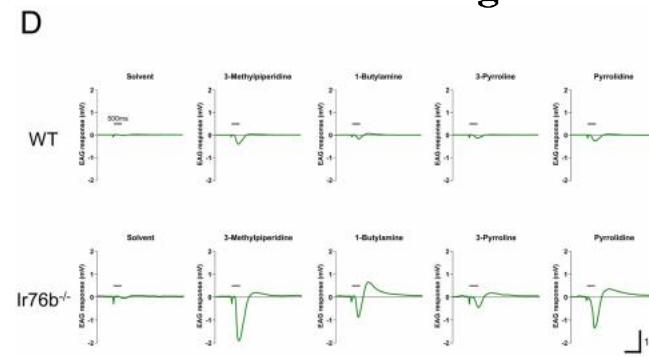
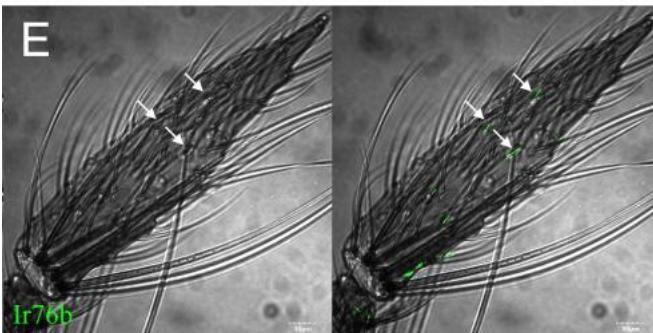
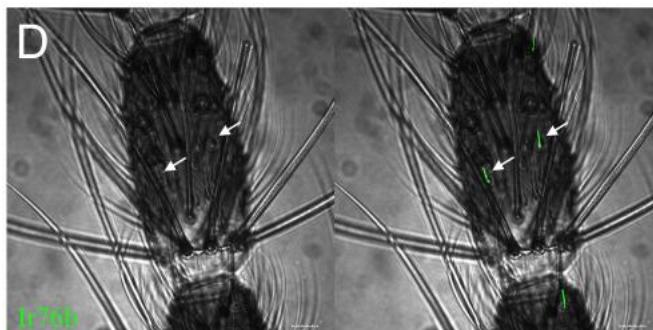
An ORN that expresses three receptors in the maxillary palp mediates the response to CO₂



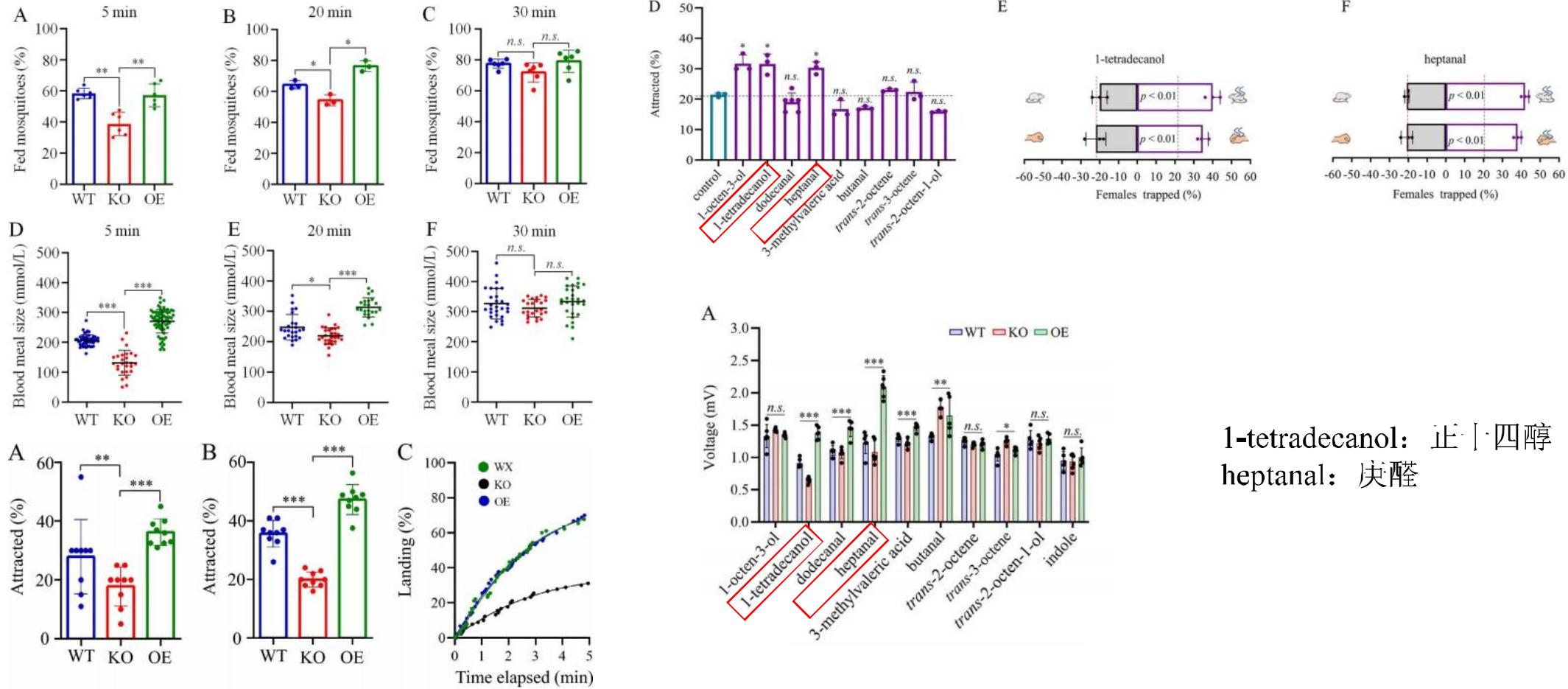
三种ORNs在下颚须的感觉器中形成固定的三联体形式

Lu T, et al. *Curr Biol.* 2007

Ir76b ionotropic coreceptor impact olfaction and blood feeding in the mosquito *Anopheles coluzzii*

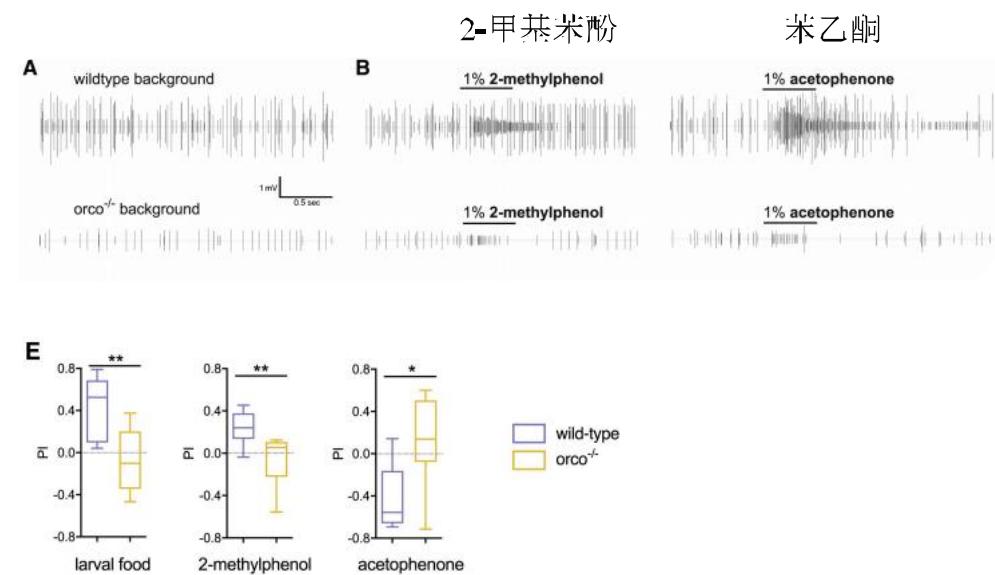
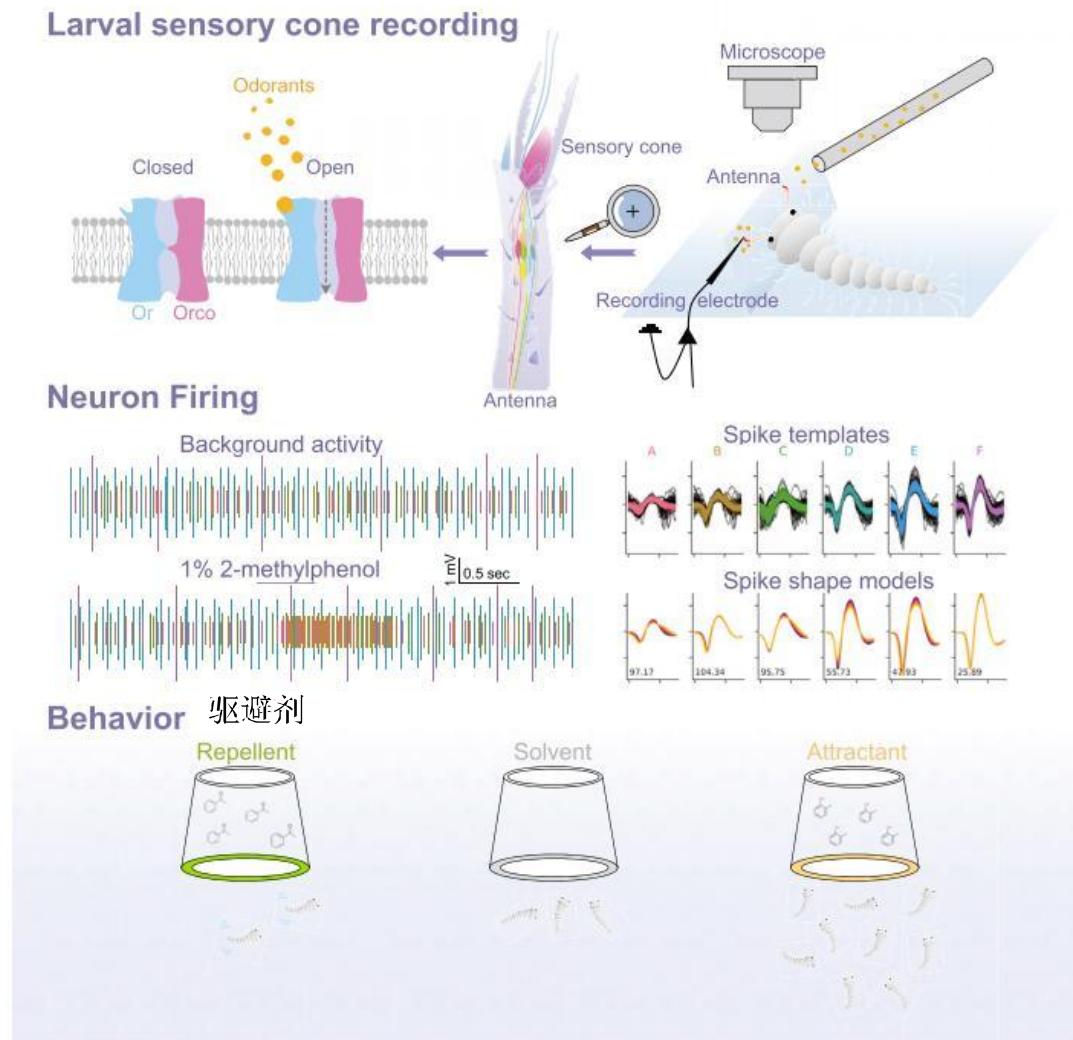


AsOBP1 is required for host seeking in the *Anopheles sinensis*



1-tetradecanol: 正十四醇
heptanal: 庚醛

Neuronal and behavioral responses to volatiles from aquatic habitats are Orco-mediated in mosquito larvae



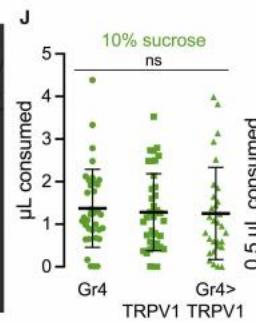
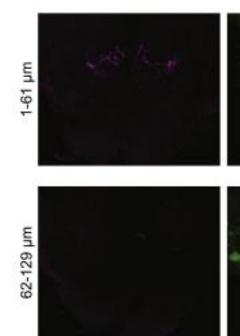
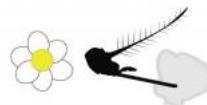
Gustation

Gustatory sensory discrimination of blood and floral nectar by *Aedes aegypti*

H Blood-feeding assay



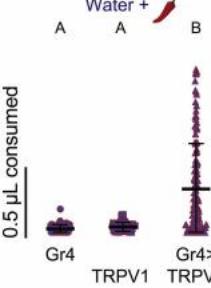
Nectar-feeding assay



Nectar-feeding assay

Water

ns



Blood-feeding assay

Water

ns

Water + chili pepper

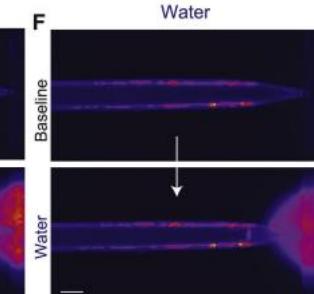
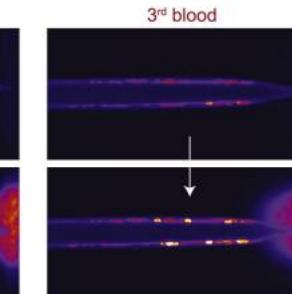
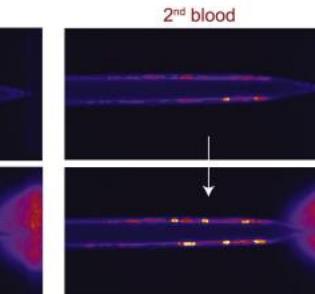
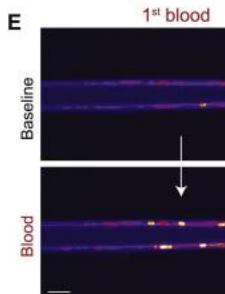
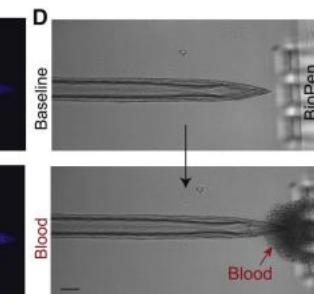
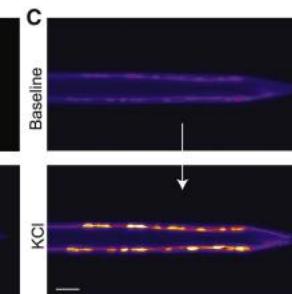
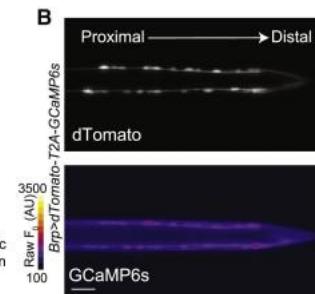
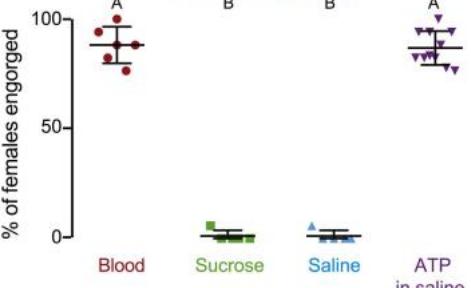
A

Water + chili pepper

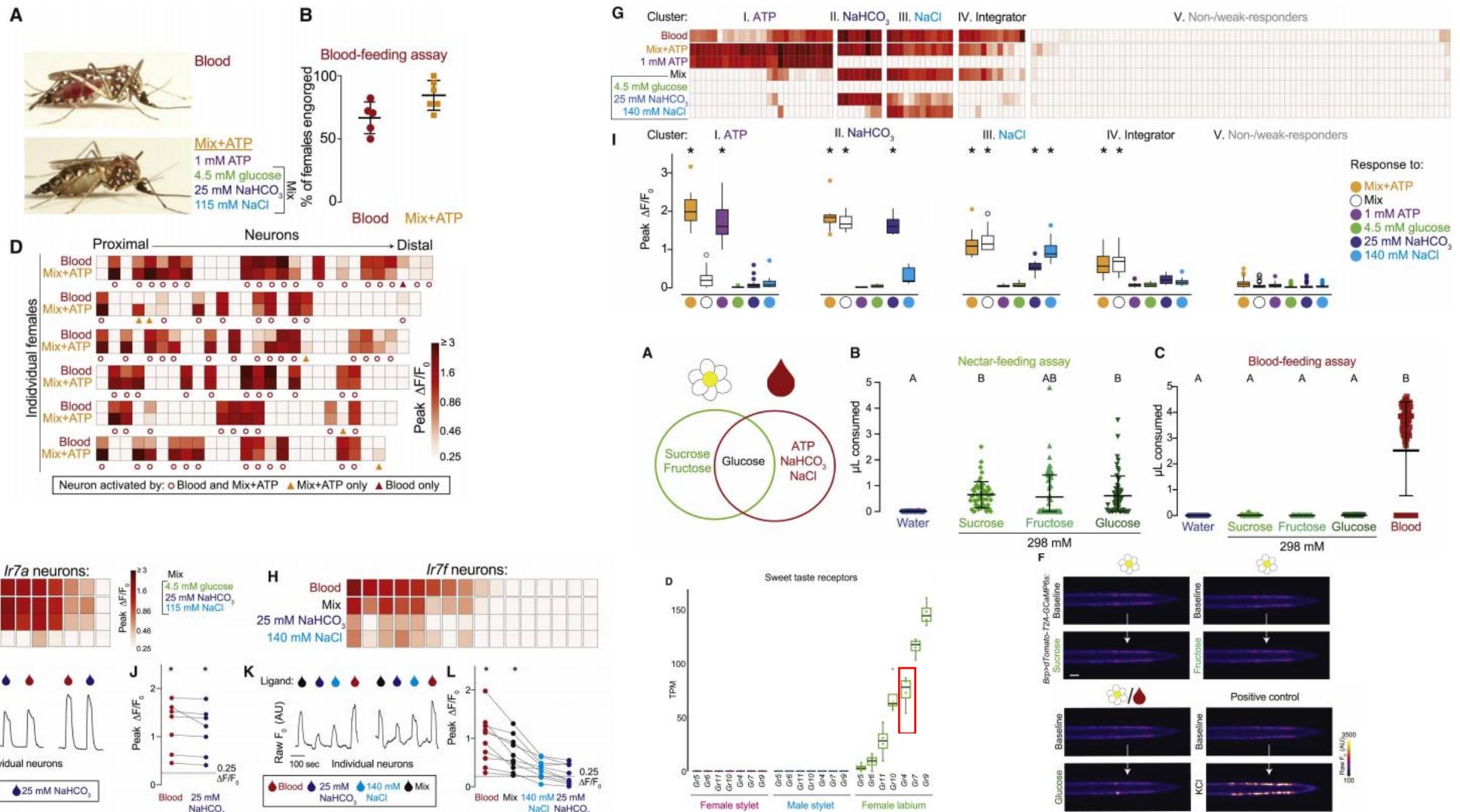
B

Water + chili pepper

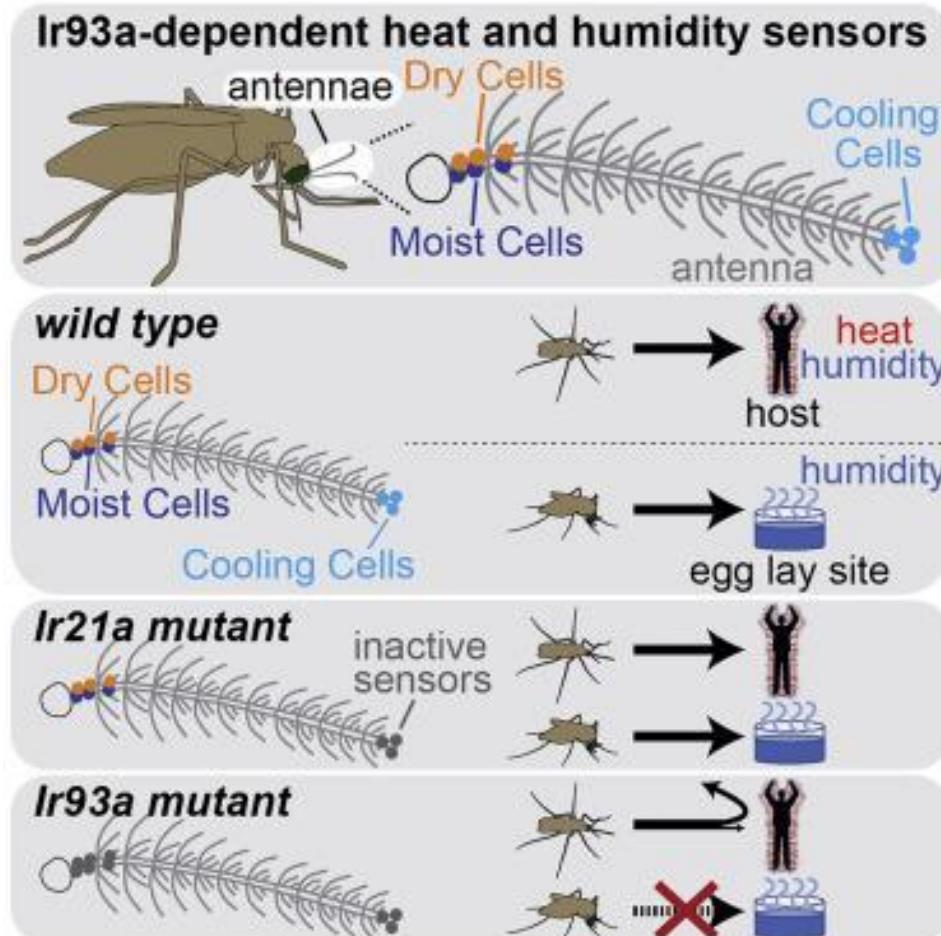
A



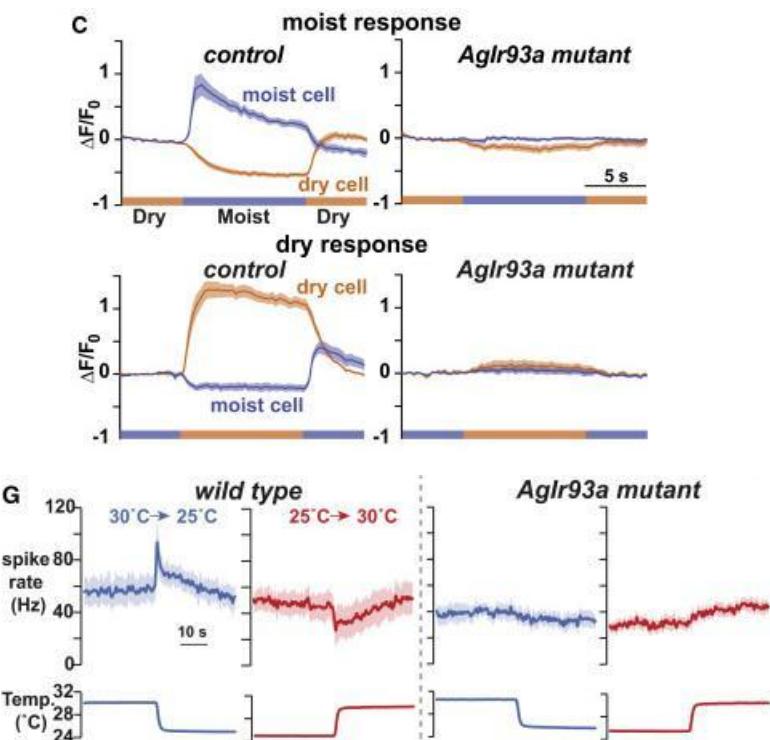
Jové V, et al. *Neuron*. 2020



• Environmental Conditions

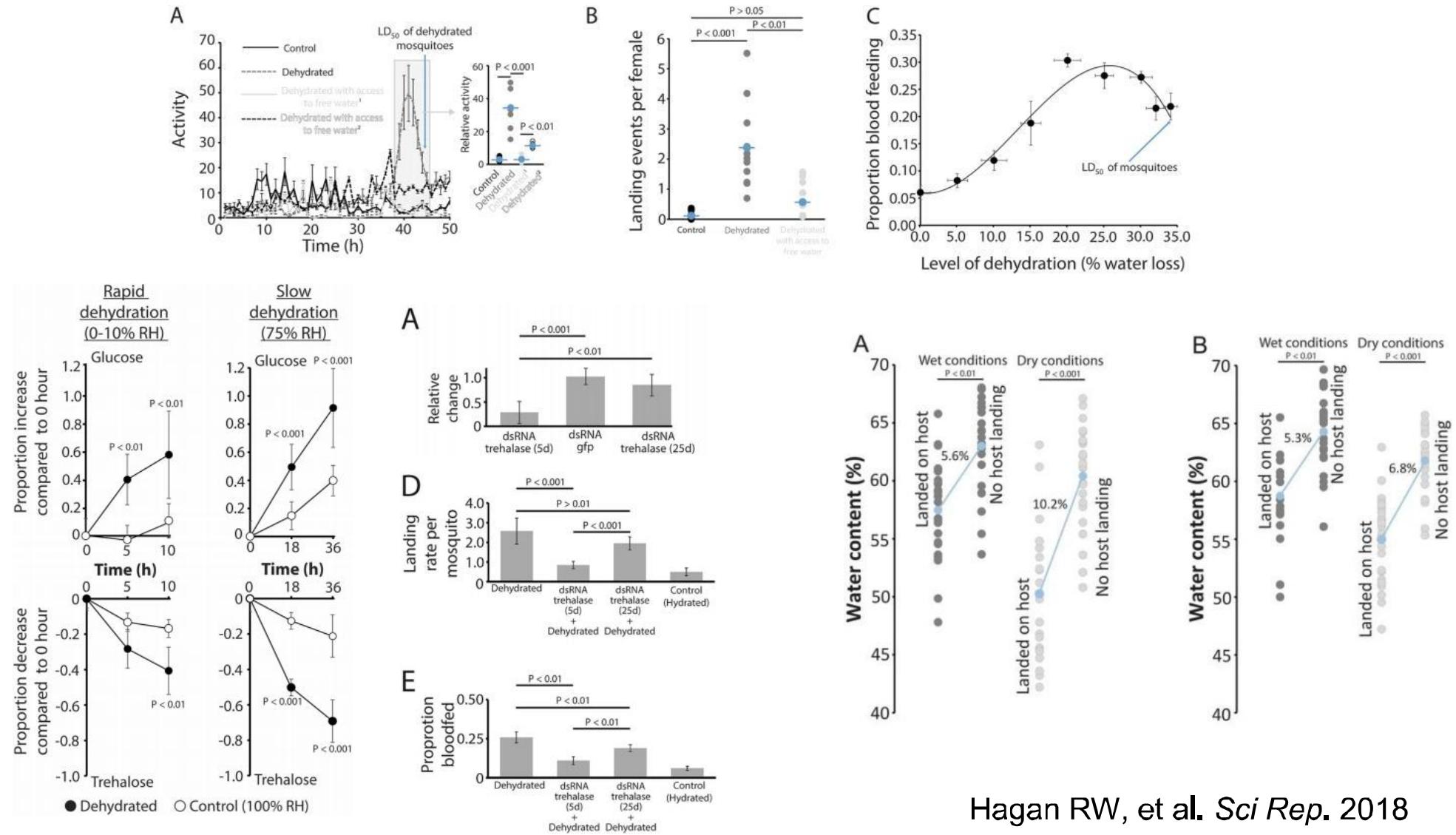


AgI93a mediates hygrosensation and thermosensation



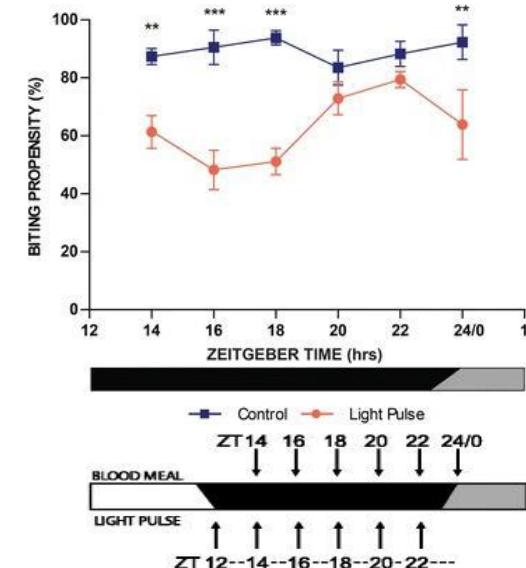
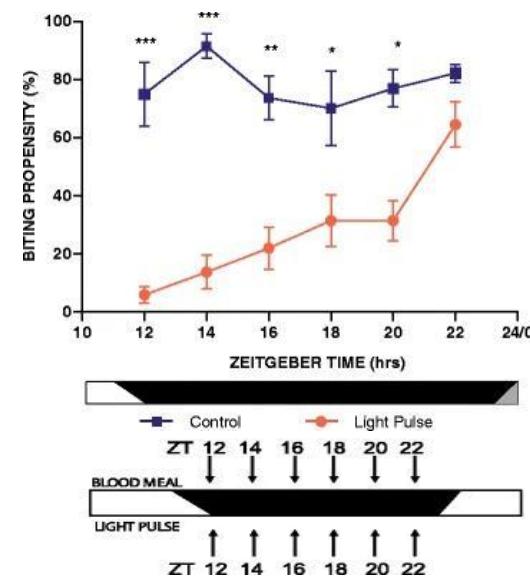
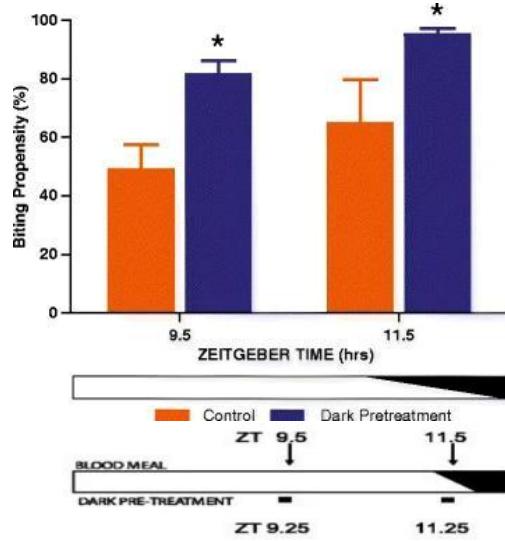
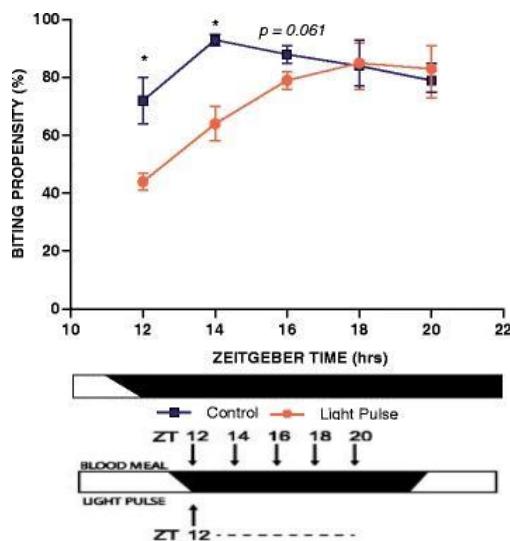
Laursen WJ, et al. *Neuron*. 2023

Dehydration prompts increased activity and blood feeding by mosquitoes



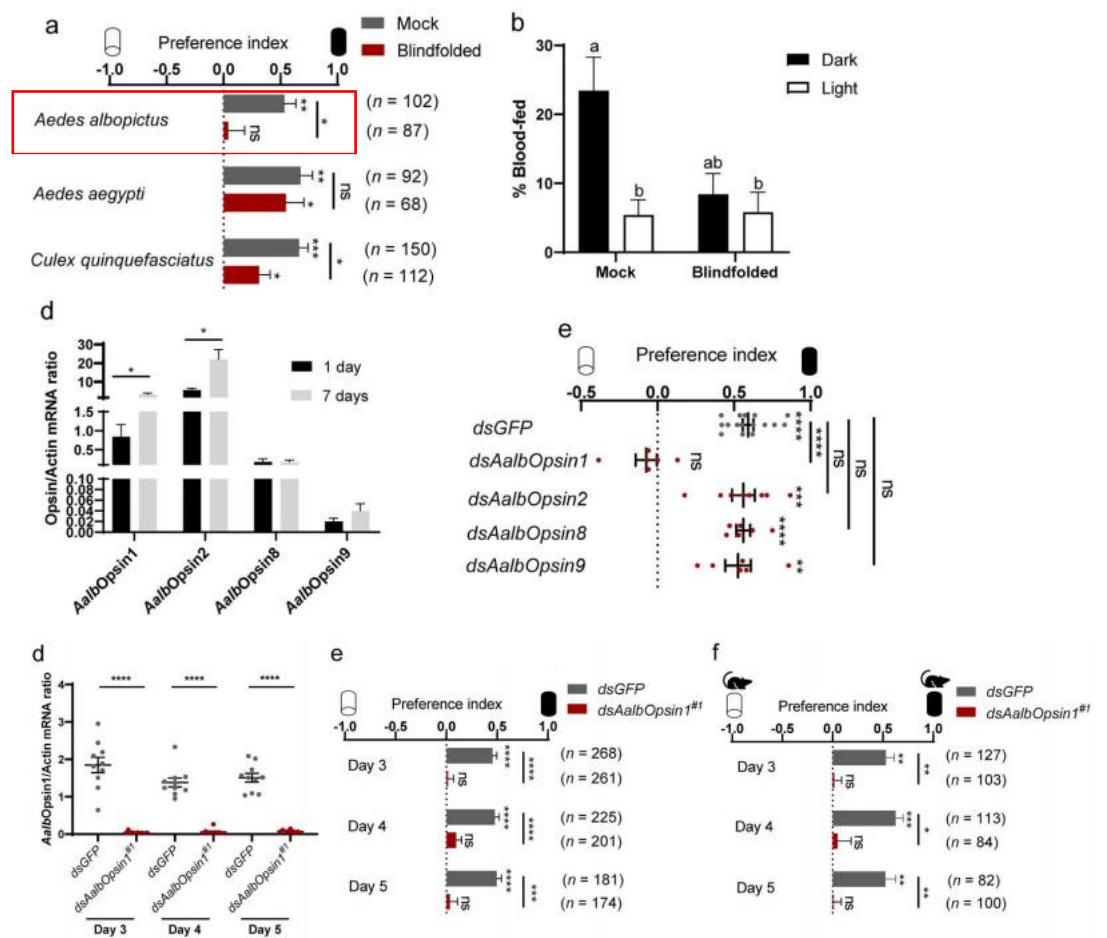
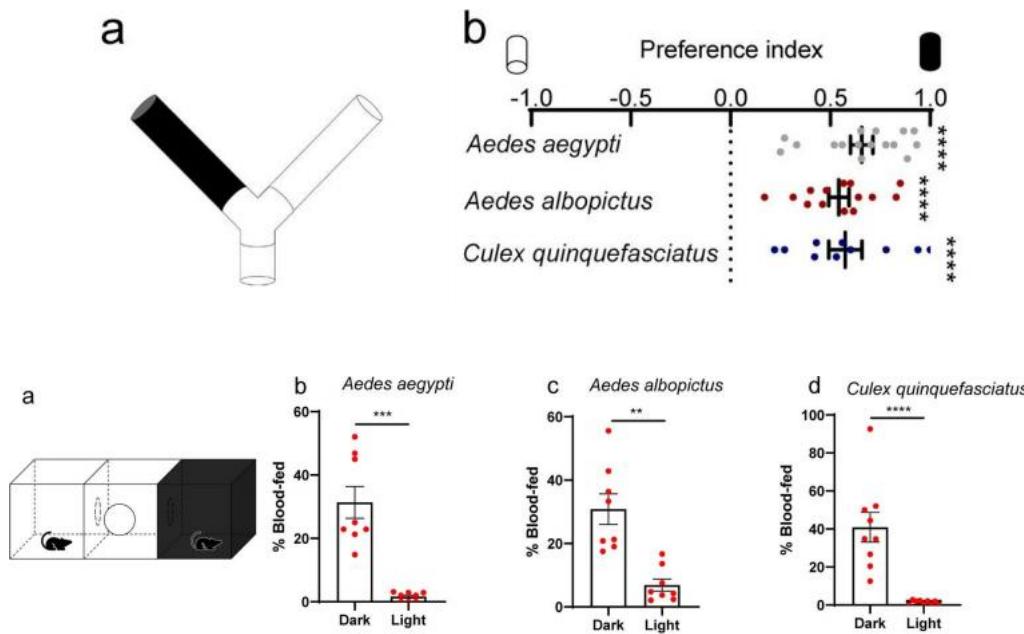
Hagan RW, et al. *Sci Rep.* 2018

Biting behavior is closely tied to photoperiod



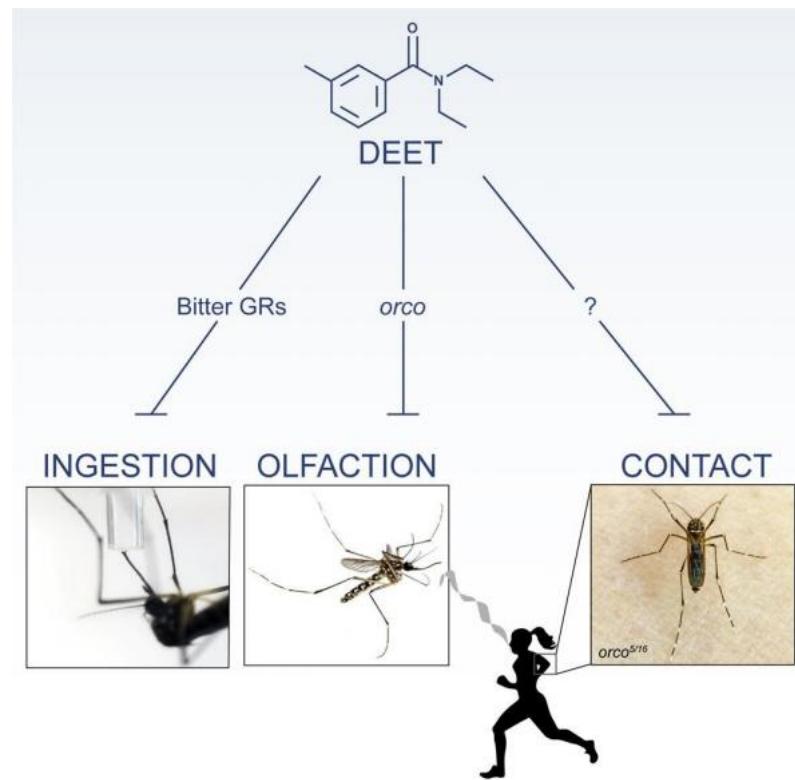
Sheppard AD, et al. *Parasit Vectors*. 2017

Opsin1 regulates light-evoked avoidance behavior in *Aedes albopictus*

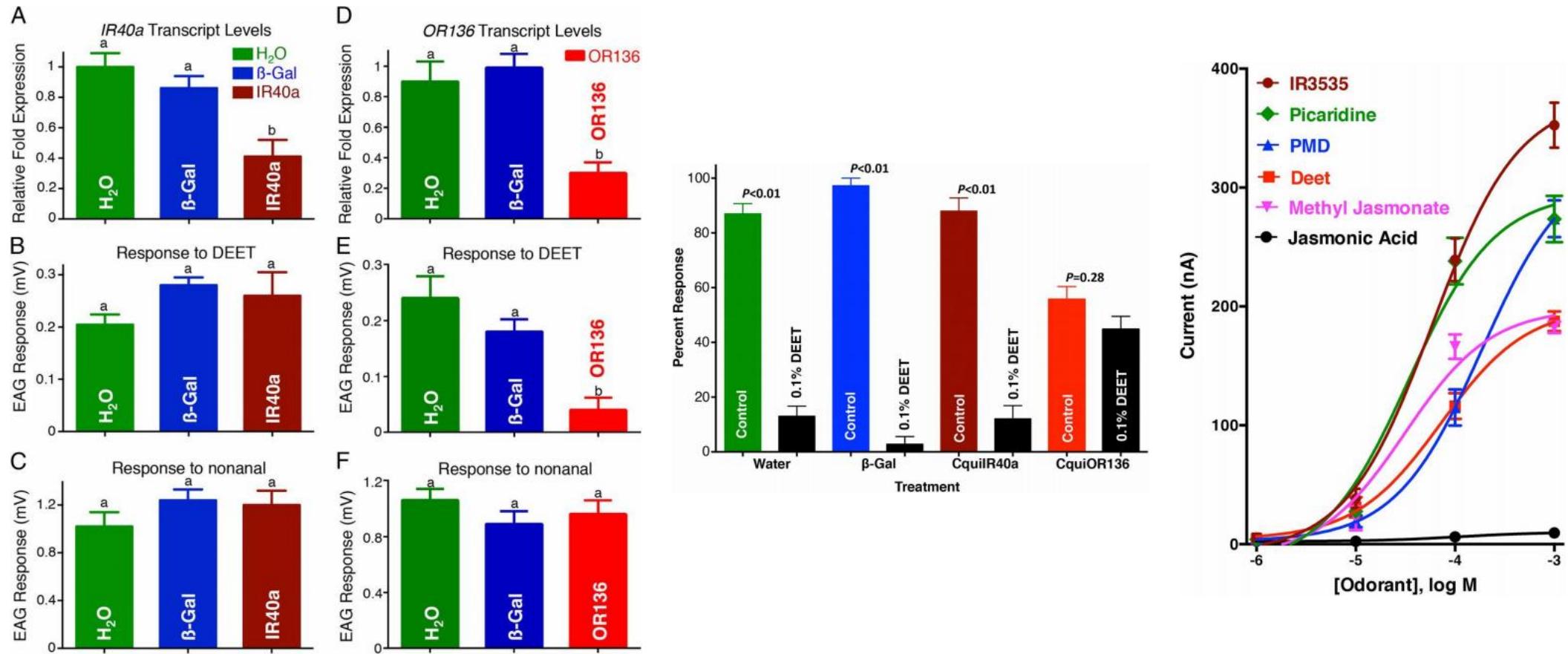


The most famous mosquito repellent - DEET

通用名称 Generic name	化学名称 Chemical name	结构式 Structure
避蚊酯、驱蚊油 DMP	邻苯二甲酸二甲酯	<chem>O=C1OC(=O)c2ccccc2C1=O</chem>
避蚊酮 Indalone	2-二甲基吡喃-4-酮-6-羧酸丁酯	<chem>CC1(C)C2C(O)C(=O)C(C1)C(=O)C2C</chem>
驱蚊醇 Rutger's 612	2-乙基-1,3己二醇	<chem>CCCC(O)CC(C)C</chem>
DEET	N, N-二乙基间甲基苯甲酰胺	<chem>CN(CC(=O)c1ccc(cc1)C)C</chem>
DEPA	N, N-二乙基苯乙酰胺	<chem>CC(=O)c1ccccc1CN(CC)C</chem>
R202	环烯酰胺	<chem>CC(=O)N(CC)c1ccccc1</chem>
依默宁 IR3535	3-(N-丁基-N-乙酰基)氨基丙酸乙酯	<chem>CC(=O)N(CC)C(C)CCN(CC(=O)OCC)C</chem>
KBR3023 Picaridin	2-羟乙基-1-哌啶酸-2-仲丁酯	<chem>CC(C)C(O)C(=O)N1CCCCC1</chem>



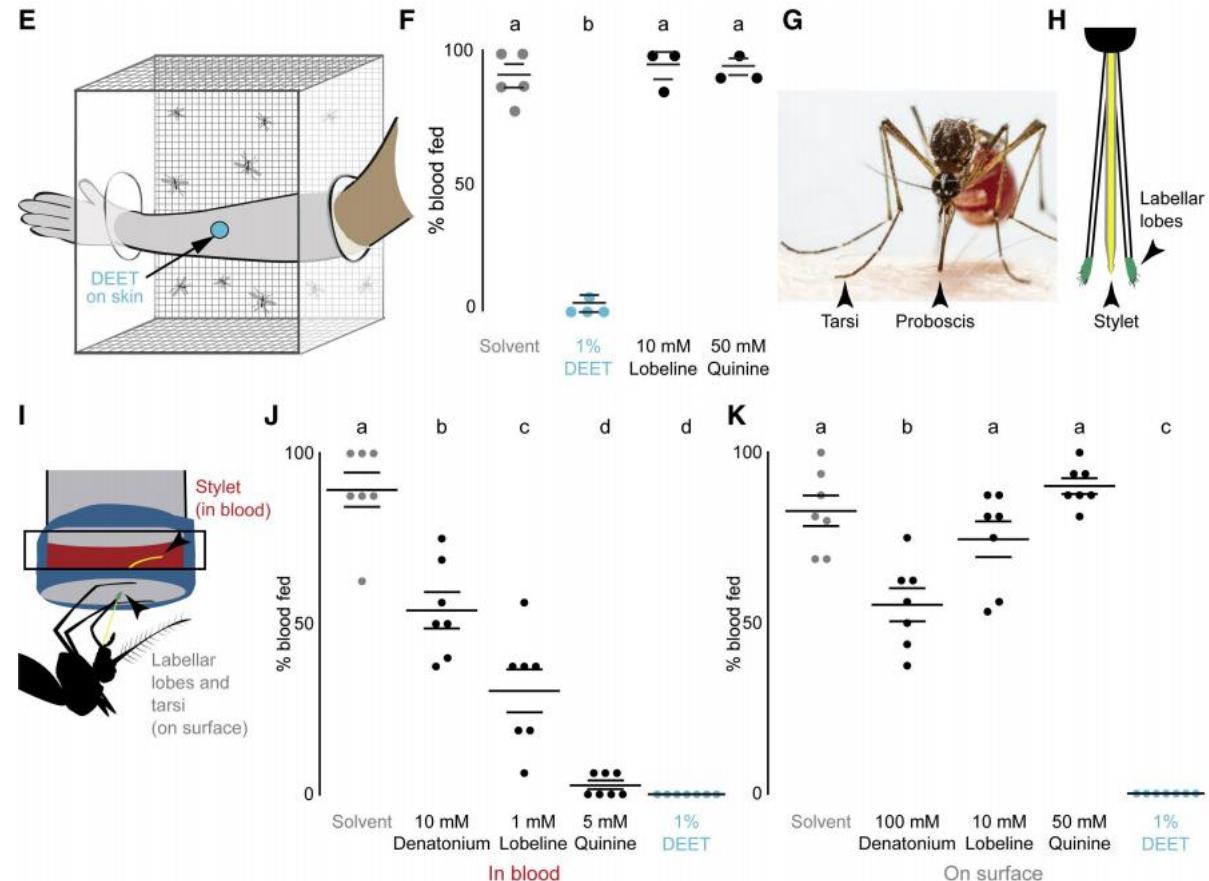
Mosquito odorant receptor for DEET and methyl jasmonate—CquiOR136



nonanal: 壬醛, 对照气味

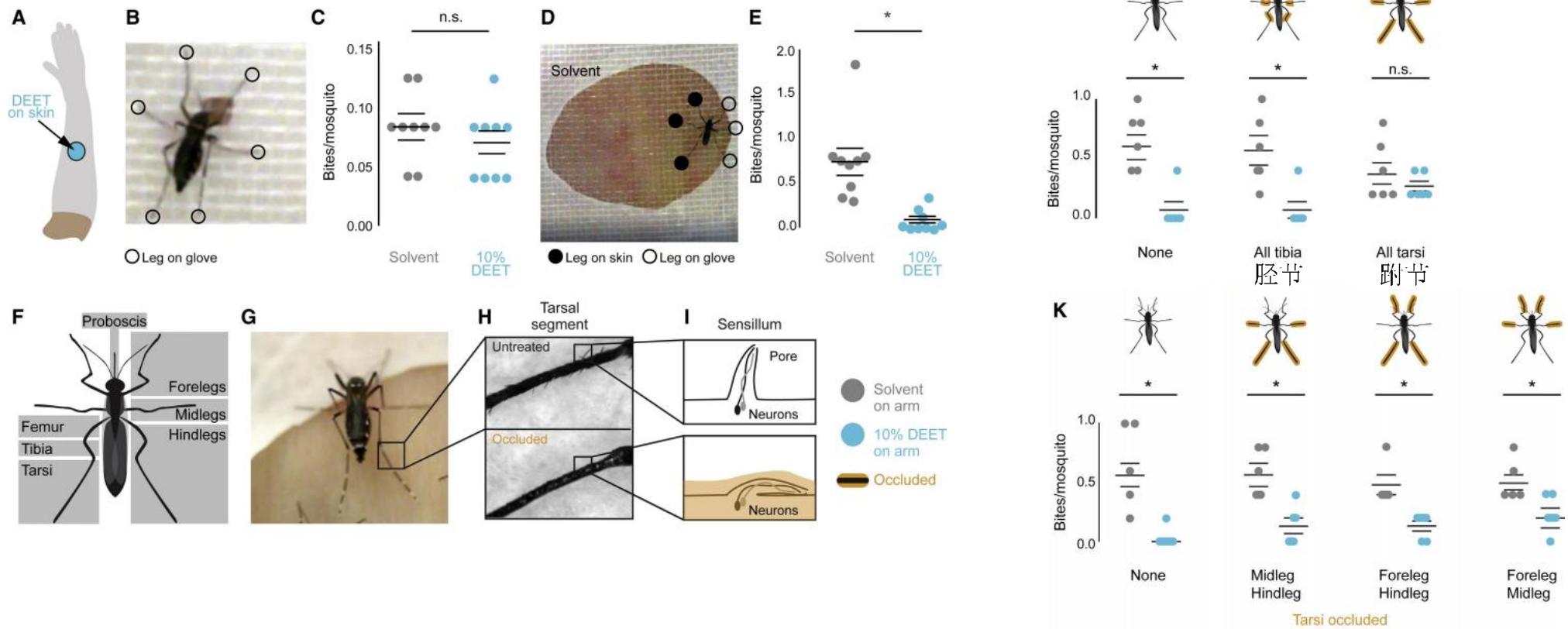
Xu PX, et al. PNAS. 2014

Aedes aegypti mosquitoes use their legs to sense DEET on contact



使用 $orco^{5/16}$ 突变蚊子来消除避蚊胺的嗅觉影响

Dennis EJ, et al. *Curr Biol.* 2019



Summary

- Mosquitoes perceive their host and changes in the external environment through multiple sensory cues.
vision, olfaction, gustation
- Disruption of any of these sensory cues could impair the mosquito's host-seeking and blood-feeding behavior.
- Gustatory neurons on the mosquito's stylet distinguish between two food components: blood and nectar.
- DEET is rejected by mosquitoes via the non-contact olfactory pathway and the contact pathway.

Thank you!