Olfaction (Chap 14)

Lecture 21

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The Chemical Senses

**Olfaction:** The sense of smell (today)

**Gustation:** The sense of taste
Odor:
A general smell sensation of a particular quality

Odorants:
- Chemical compounds
- But not every chemical is an odorant
- Most are small, volatile, and hydrophobic (don’t diffuse in water)
Is smell even relevant to humans?

our “machinery” is less sensitive than other animals (dogs, etc.)

• dogs can detect odorant concentrations 100x lower than humans (dogs: can detect 1 part-per-million, humans: 100 parts-per-million)

• but, experiments show that human receptors respond to single odorant molecules

• the difference? Dogs have ~ 1 billion receptors, humans have ~ 10 million
Conventional wisdom: humans not very good at olfaction

Bloodhound tracking a pheasant through a field

Conventional wisdom is wrong!

Human tracking a scent trail through a field

Porter et al 2007
Two nostrils sample different regions of space
single- vs. dual-nostril sniffing

- humans use info from both nostrils for scent tracking

(24% slower w/ 1 nostril)
How good is our sense of smell?

• Latest findings suggest we can detect over one trillion smells!

• We can only detect about 7.5 million colors.

(Oh really!)
Physiology of the Olfactory System
The nose

- **Olfactory cleft**: space at the back of the nose into which air flows, where the main olfactory epithelium is located.
- **Olfactory epithelium**: secretory mucosa whose primary function is to detect odorants.
Olfactory epithelium: the “retina of the nose”

Three cell types

- **Supporting cells**: Provides metabolic and physical support for the olfactory sensory neurons
- **Basal cells**: Precursor cells to olfactory sensory neurons
- **Olfactory sensory neurons (OSNs)**: The main cell type in the olfactory epithelium

  - OSNs make direct contact with physical stimulus (i.e., unlike in retina, cochlea, or skin)
Olfactory sensory neuron

- Responses are slow!
- OSN axons among the thinnest and slowest in the body
Olfactory epithelium: the “retina of the nose”

**Cilia:** Hairlike protrusions on OSN dendrites
- Have receptor sites for odorant molecules.
- Structures for olfactory signal transduction

**Olfactory receptor (OR):** The region on the cilia of OSNs where odorant molecules bind

- Takes seven or eight odor molecules binding to a receptor to initiate an action potential
Olfactory epithelium: the “retina of the nose”

**Cribiform plate**: bony structure with tiny holes (even with the eyebrows), separating the nose from the brain

- Axons from OSNs pass through the tiny holes to enter the brain
Olfactory epithelium: the “retina of the nose”

**Mitral cells:** The main projective output neurons in the olfactory bulb

**Glomeruli:** Spherical conglomerates containing the incoming axons of the OSNs

- Each OSN converges on two glomeruli
Sensory pathway for olfactory system
**Olfactory bulb**: The blueberry-sized extension of the brain just above the nose, where olfactory information is first processed

- There are two olfactory bulbs, one in each brain hemisphere, corresponding to the left and right nostrils.
Primary olfactory cortex: cortical area where olfactory information is first processed.

Limbic system:
- Involved in many aspects of emotion and memory
- Olfaction is unique for its direct connection to limbic system
Why Olfaction is Weird

- direct connection to cortex
- many subsequent cortical connections to midbrain
- also, no “smell-o-topic” maps: no topography
Olfactory Physiology

**Anosmia**: The total inability to smell, most often resulting from sinus illness or head trauma

- A hard blow to the front of the head can cause the cribriform plate to be jarred back or fractured, slicing off the fragile olfactory neurons
- Anosmia causes a profound loss of taste as well as smell

Essay: “The Miseries of losing one’s sense of smell”
http://www.slate.com/id/2195018/
Genetic basis of olfactory receptors:

• Buck and Axel (1991) showed that genome contains about 1000 different olfactory receptor genes; each codes for a single type of OR

• All mammals have pretty much the same 1000 genes.
• However, some genes are non-functional “pseudogenes”
  ▪ Dogs and mice: About 20% are pseudogenes
  ▪ Humans: Between 60% and 70% are pseudogenes

Each person has a different number of pseudogenes, resulting in individual differences in sensitivity to smells
Evolutionary trade-off between vision and olfaction

% pseudogenes

Old World Primates

New World Primates

species (color-coded by family)

Gilad et al, PLoS 2004
Evolutionary trade-off between vision and olfaction

% pseudogenes

species (color-coded by family)

Gilad et al 2004
Black arrows indicate on which lineages the acquisition of full trichromatic color vision occurred.

Red lines show lineages with a high proportion of OR pseudogenes.

Gilad et al 2004
shape-pattern theory of olfactory perception

- scent percept depends on fit between OR shape and odorant shape
Theories of olfactory perception:

• **Shape-pattern theory**: The current dominant theory.
  
  - The binding pattern of odorants in the olfactory epithelium produces specific firing patterns of neurons in the olfactory bulb, which then determine the particular scent we perceive.

• **Vibration theory**: now defunct.
  
  - Proposes that every perceived smell has a different vibrational frequency, and that molecules that produce the same vibrational frequencies will smell the same.

(or is it? See this bizarre TED talk: [https://www.ted.com/talks/luca_turin_on_the_science_of_scent](https://www.ted.com/talks/luca_turin_on_the_science_of_scent))
Study of **stereoisomers**

- Molecules that are mirror-image rotations of one another; although they contain the same atoms, they can smell completely different
- Vibration theory cannot explain this phenomenon

Theory that molecules with similar vibration frequencies should smell similarly

(a) \textit{d-carvone} \hspace{1cm} (b) \textit{l-carvone}

Smells of caraway \hspace{1cm} Smells of spearmint
The importance of patterns

• *How can we detect so many different scents if our genes only code for about 1000 olfactory receptors?*

• We can detect pattern of activity across many receptor types

• Intensity of odorant changes which receptors are activated (Weak concentrations of an odorant may not smell the same as strong concentrations of it!)

• Specific time-order of activation of OR receptors is important
Bi-nostral smelling: why have two nostrils?

“The world smells different to each nostril”
Sobel et al, Nature 2000

Background
1. Airflow is greater into one nostril than the other, due to slight swelling that obstructs airflow.
2. Switches nostrils several times per hour.

Q: What are the consequences for olfaction?
Bi-nostral smelling: why have two nostrils?

Background
1. Airflow is greater into one nostril than the other, due to slight swelling that obstructs airflow.
2. Switches nostrils several times per hour.

Obstructed nostril
(swollen turbinates)

un-obstructed nostril
(relaxed turbinates)
• Odorants sorb across nasal mucosa at different rates

- “high-sorption” odorant – induces small response when airflow is low, and large one when airflow is high
- “low-sorption” odorant – large response when airflow is low; small

**Finding:** odorants do indeed smell different in nostrils, depending on the air flow and sorption of the odorant!
Olfactory Psychophysics

• How much stimulation is required before we perceive something to be there?

• Olfactory detection thresholds: Depend on several factors
  ▪ Women: Generally lower thresholds than men, especially during ovulatory period of menstrual cycles, (but sensitivity is not heightened during pregnancy)
  ▪ Professional perfumers and wine tasters can distinguish up to 100,000 odors

(or is it 1 trillion???)
And also:

- Age: By 85, 50% of population is effectively anosmic

(like those high-pitched noises, enjoy smelling while you still can!)
Olfactory Hedonics

Odor hedonics: the “liking” dimension of odor perception - measured with scales of pleasantness, familiarity, and intensity

Familiarity and intensity:

• Pleasantness: obvious
• Familiarity: tend to like odors we’ve smelled before
• Intensity: more complicated relationship with odor liking
Odorants: Pleasantness vs. intensity

(a) | (b)
---|---
**Phenyl ethyl alcohol** | **Trimethylamine**

- **P** Pleasant
- **N** Neutral
- **U** Unpleasant

Inverted U-shaped function | Linearly decreasing function
Olfactory Hedonics

Nature or nurture?

• Long-standing debate: innate vs. learned
• **verdict**: almost completely “nurture”
• infants: not put off by sweat or feces; don’t discriminate banana from smell of rancid food
• Cross-cultural data support associative learning

• Wintergreen study (Moncrief, 1966)
  - Americans like it.
  - English rated it the most unpleasant of many odors (used in medicine)

• US Army: tried to develop stink bomb for crowd dispersal: couldn’t find a smell that was universally disgusting (including “US Army Issue Latrine Scent”)

Japanese and American people have very different tastes in food

Cheese
• disgusting to most Japanese

Natto
• fermented soybeans; Japanese breakfast food
Olfactory Hedonics

• Evolutionary argument: *generalists* (like us, and roaches) don’t need innate smell aversions to predators

• **learned taste aversion**: Avoidance of a flavor after it has been paired with gastric illness.
  - finding: from the smell, not the taste (Bartoshuk 1990)
Olfaction and memory

Q: are odors really the best cues to memories?

- Memories triggered by odor cues are distinctive in their emotionality
- But *not* (it turns out) more accurate

The smell, sight, sound, feel, and verbal label of popcorn elicit memories equivalent in terms of accuracy but not emotion.
Olfaction summary

- odors, odorants
- scent tracking, binostril smelling (2 reasons)
- olfactory cleft, olfactory epithelium
- Olfactory Receptors (ORs), located on cilia
- Olfactory Sensory Neurons (ORNs)
- cribiform plate, glomeruli, mitral cells, olfactory bulb, primary olfactory cortex, limbic system
- anosmia
- pseudogenes and trichromatic color vision
- shape-pattern theory
- olfactory hedonics, learned taste aversion
- olfaction and memory
- pheromones / chemosignals & VNO (in book)