



Acupuncture, the Vagus, & Modern Interpretations

Michigan Osteopathic Association

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Mathew Kulas, L.Ac. – Speaker
I have no relevant financial relationships to disclose



Intro

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Intro

- Licensed Acupuncturist and Chinese Herbal Practitioner
- Masters of Acupuncture from The Academy for Five Element Acupuncture, Gainesville, FL
- Transitional Doctorate of Acupuncture from Pacific College of Health Sciences, San Diego
- Been with HF for 7+ years
- Passionate about helping pts w/ pain, trauma, sleep, tinnitus, cancer, whole body wellness, and recently dysautonomia conditions
 - Auricular vagal stimulation + diaphragmatic breath work



Modalities

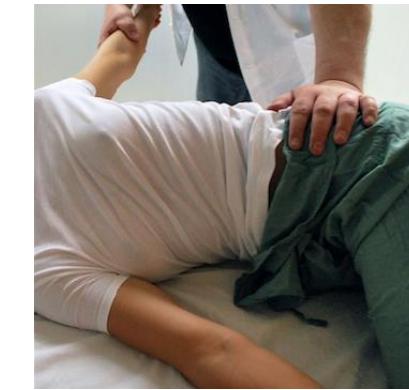
Herbal Medicine



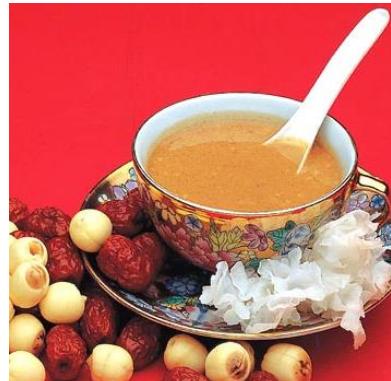
Acupuncture



Manual Therapies



Diet / Lifestyle



Exercise / Breathing



Objectives

- What is Acupuncture?
 - Classical perspective
 - Modern mechanisms
- Vagal mechanism and importance
- Integrative & translational approach

Goals

- Acupuncture is *REAL* science

Indications

National Center for Complementary and Integrative Health (NCCIH)

- Back/neck pain
- OA
- Headache
- Myofascial pain
- Sciatica
- Post-op pain
- Cancer pain
- Irritable Bowel
- Chronic Prostatitis / Pelvic pain
- Fibromyalgia
- Seasonal Allergies
- Urinary Incontinence
- Nausea / Vomiting
- Asthma
- Depression
- Smoking Cessation
- Carpal Tunnel
- Menopausal Hot Flashes

Best referrals

- Refractory to traditional therapies, such as:
 - PT – even if px have done dry needling
 - Medications
 - Lifestyle adjustments
 - When you suspect a psychosomatic cause or *barrier*
 - MVA, Veterans, etc
 - Acupuncture is **non-verbal** healing
- Optimizing overall health
 - Athletes, increase ADLs and QOL
- When pragmatic approaches are needed

What is Acupuncture?

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Classical Philosophy

- QI = energy
- Lots of different kinds of qi and ways to cultivate:
 - Air
 - Food
 - Water
 - Exercise
 - Meditation
 - Sleep
 - ATP
 - Nervous system action potentials
 - Chemical: digestive enzymes, bile
- **Yuan qi, zhong qi, kidney qi, wei qi, etc**
- **TRANSFORMATIVE**
- **CULTIVATED**
- **OPTIMIZATION**
- **SALUTOGENIC**



Acupuncture & Moxibustion

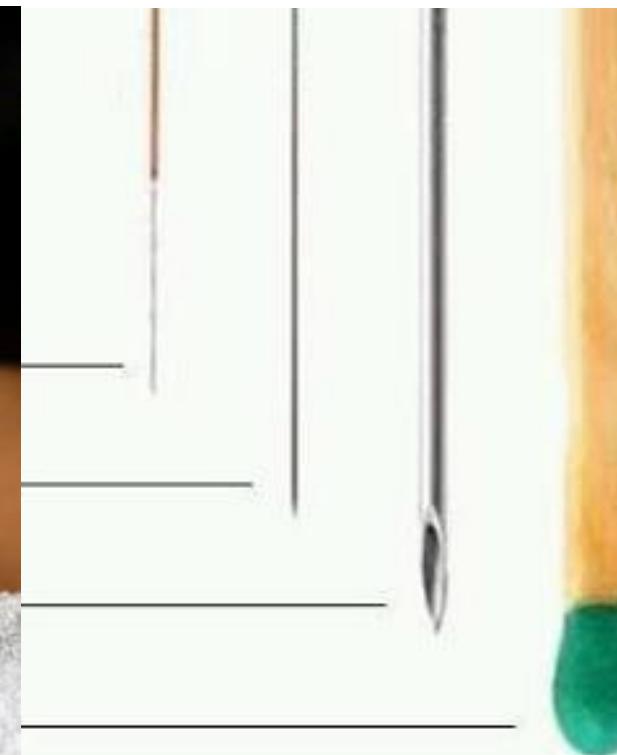
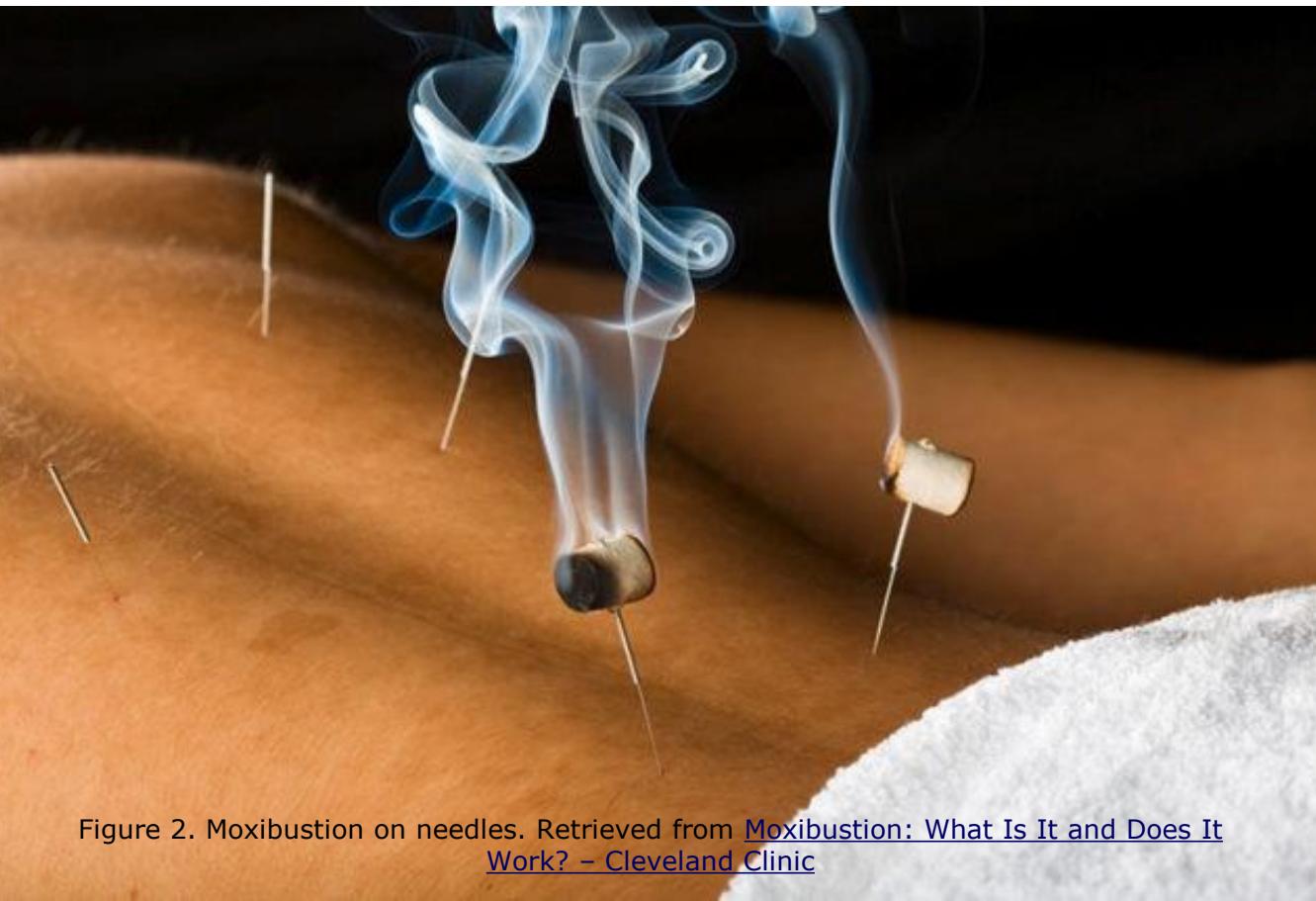
针灸

zhēn jiǔ

acupuncture and moxibustion

to give or have acupuncture and moxibustion

Trad. 针灸



How tiny
is an
acupuncture
needle?

Figure 2. Moxibustion on needles. Retrieved from [Moxibustion: What Is It and Does It Work? – Cleveland Clinic](#)

Figure 1. Acupuncture size comparison. Retrieved from [Acupuncture FAQs | Bret Idenhauer | Acupuncture, Cupping, Pain Management, Sports | Norspring Cryotherapy, Chattanooga, TN; \(instituteforacupuncture.com\)](#).

Acupuncture

- 12 main meridians and organs/officials:
 - Heart & Small Intestine
 - Bladder & Kidneys
 - Pericardium & Triple Burner
 - Gallbladder & Liver
 - Lung & Large Intestine
 - Stomach & Spleen
 - 2 foundational meridians:
 - Governing and Conception Vessels
 - Meridians give us access to pools, vortexes, or POINTS to manipulate qi

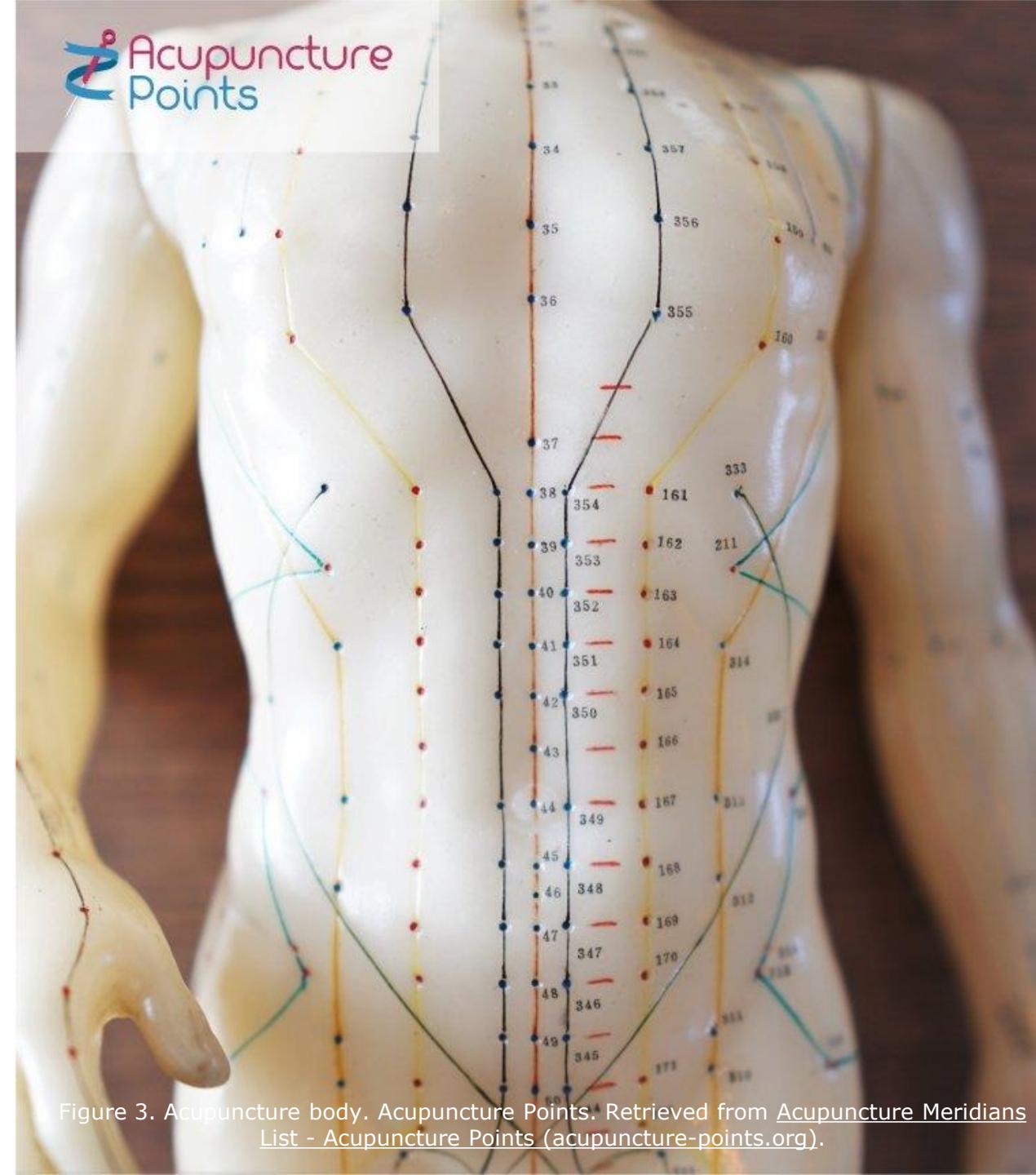
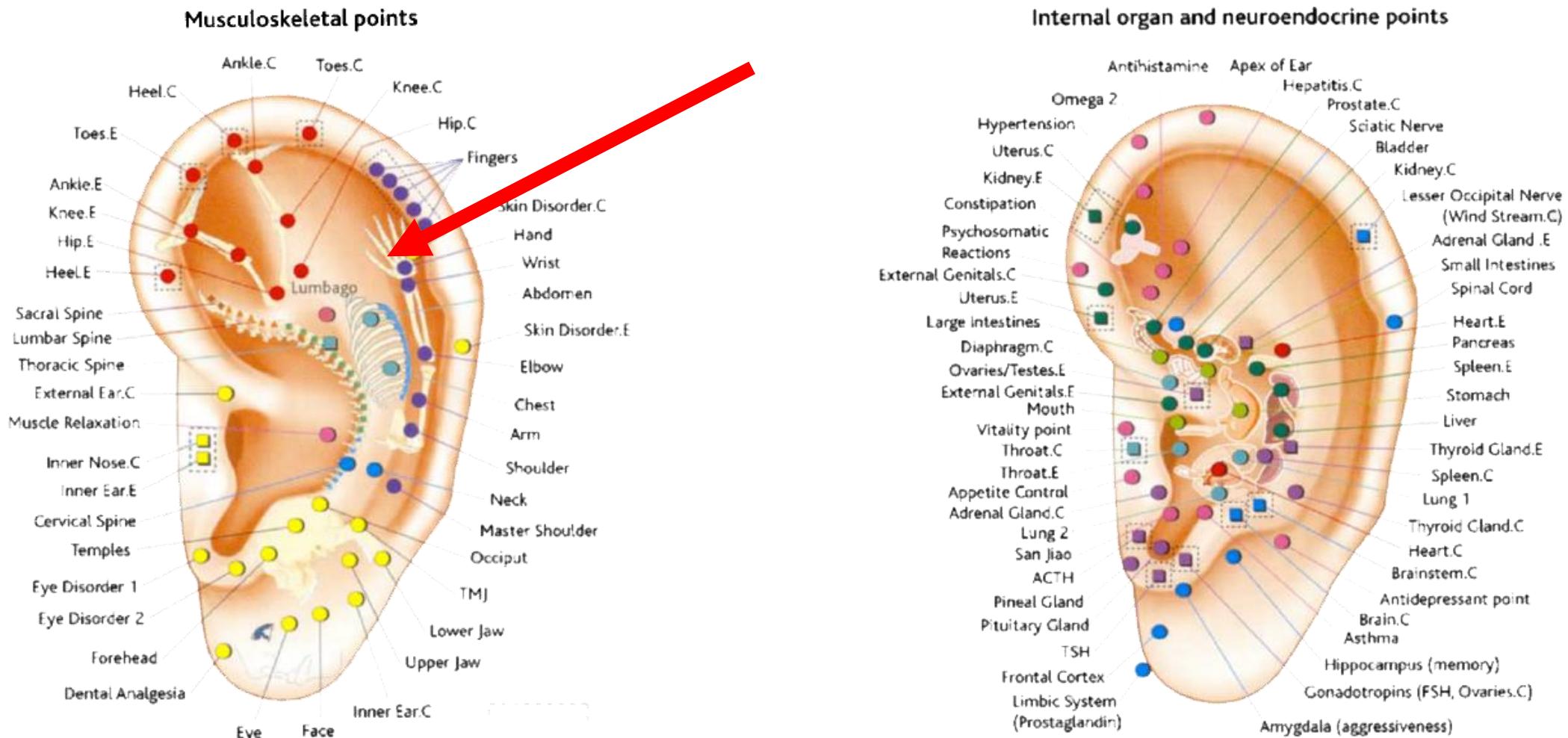


Figure 3. Acupuncture body. Acupuncture Points. Retrieved from [Acupuncture Meridians List - Acupuncture Points \(acupuncture-points.org\).](http://Acupuncture Meridians List - Acupuncture Points (acupuncture-points.org).)

Auricular Acupuncture



Variations, Synonyms, & Adjunctive therapies

- Traditional Chinese Medicine (TCM)
- Five Element
- Korean hand, Japanese
- Medical Acupuncture
- **Dry needling, trigger point, motor point**
- **Intramuscular stimulation**
- Guasha/Graston, Cupping, tuina massage
- 8 extraordinary meridians
- Microsystems (auricular, fascial, scalp)
- Medical Qi gong

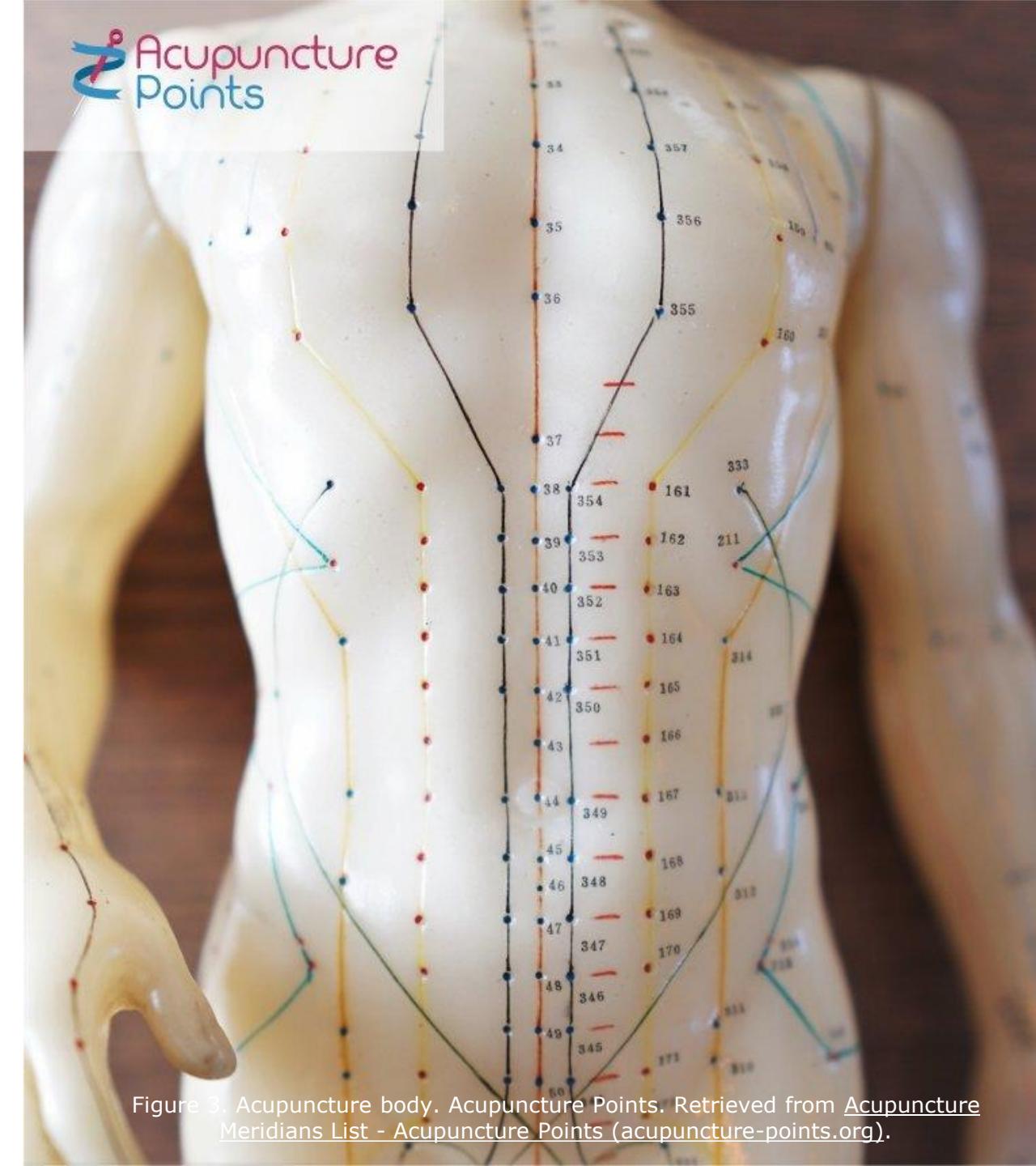


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Treatment strategies

- Strategies:
 - Distal needling
 - Local needling
 - Auricular points
 - Acupressure
 - Electric stimulation vs. manual
 - Specific protocols, point combos
 - Pragmatic:
 - Lifestyle, supplements, herbs
 - Recommendations:
 - 6-10 treatments minimum to start
 - 1-2x per week
 - High responders, low responders, middle

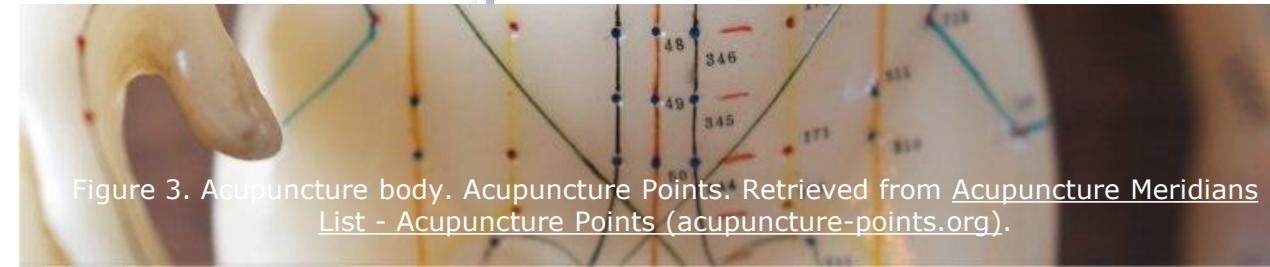
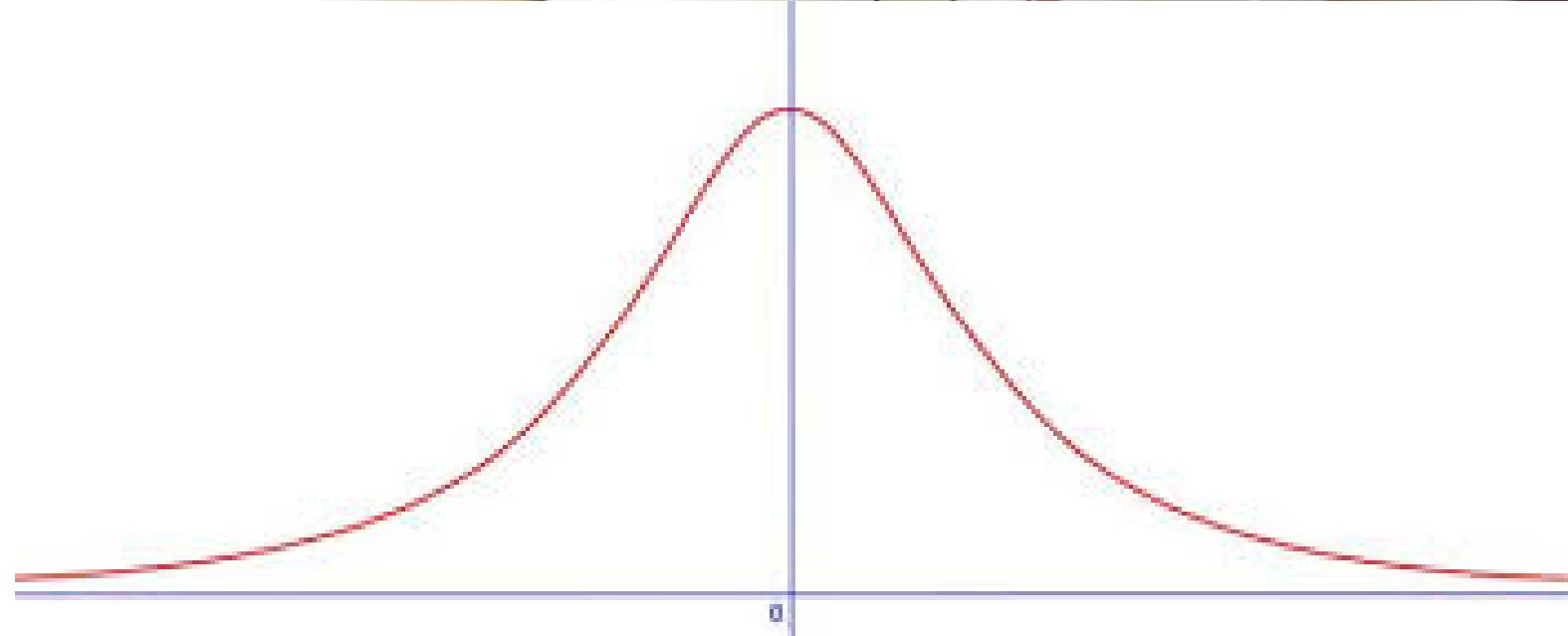
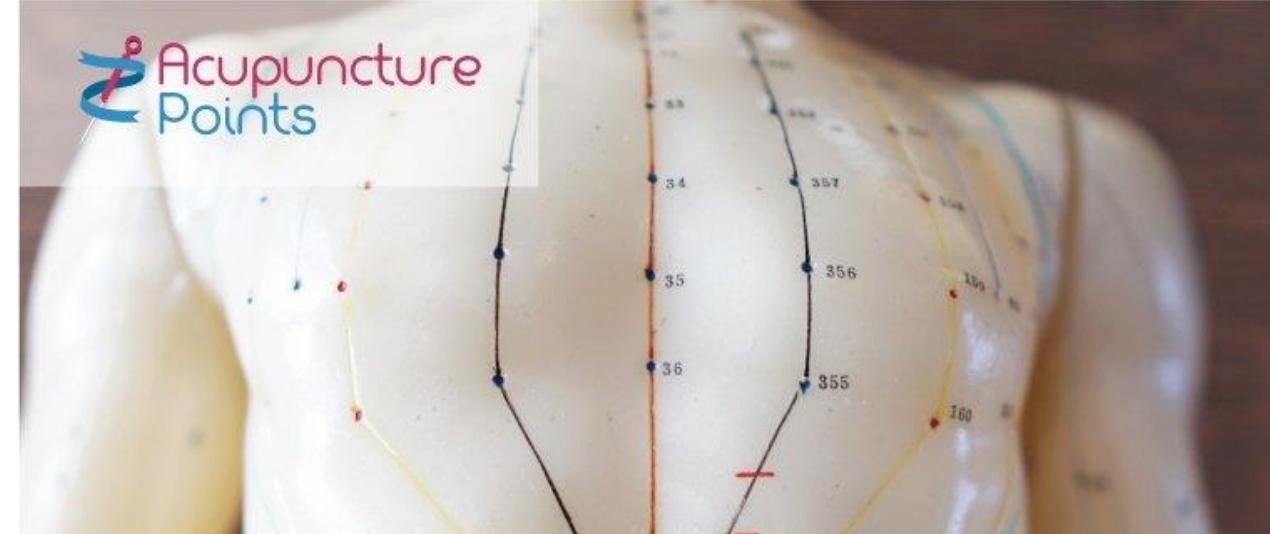


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Diagnostics

- Tongue
- Pulse
- “10 questions”
- Physical assessment
- Kidney yin deficiency
- Qi deficiency & phlegm
- Liver wind
- Running piglet syndrome
- Normal tongue

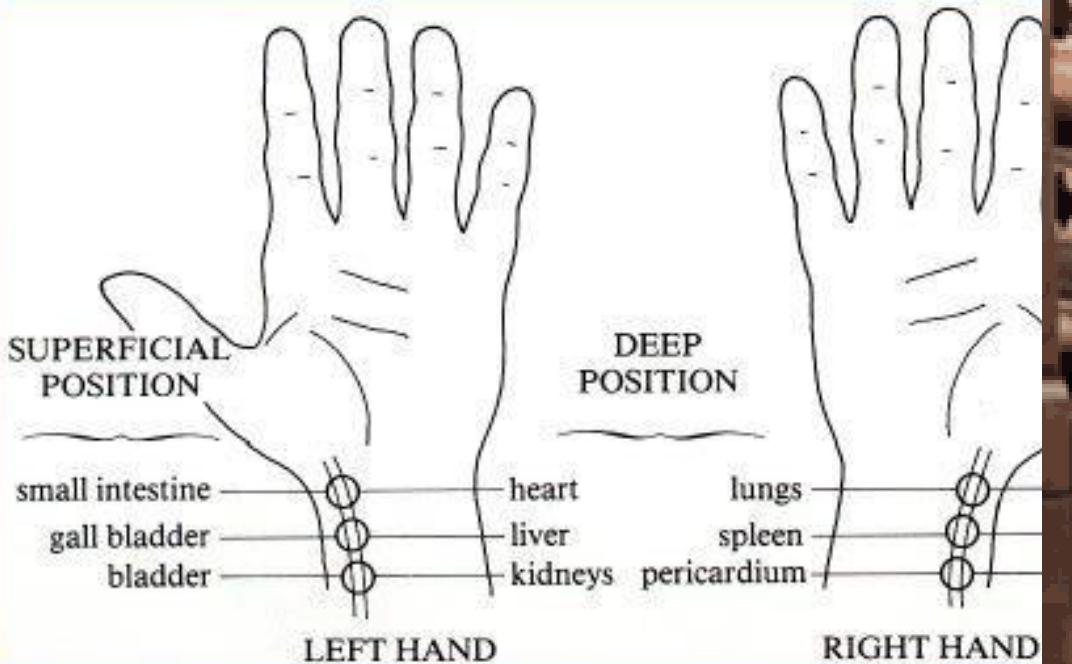


Figure 6. Pulse positions
Chinese pulse diagnosis
Chinese pulses|Electro-
(doctoracupuncture.co.uk).



But how does
acupuncture *REALLY*
work!?

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Modern Interpretations

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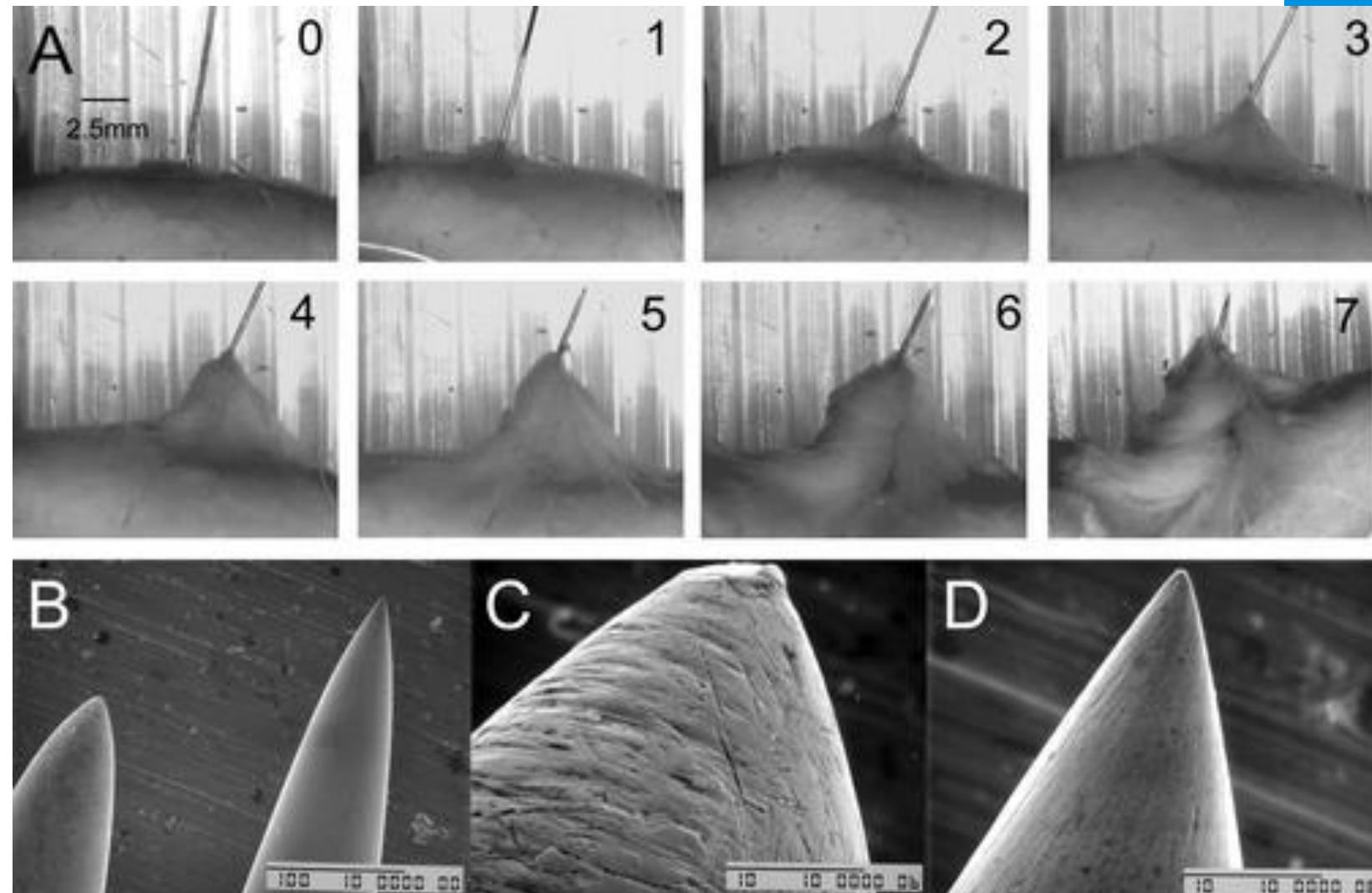
Pluripotent Mechanisms

Gate Control Theory	Endocrine Theory	Endorphin Theory	Neuro-immune Theory	Circulatory Theory
<p>It generates a competing stimulus which blocks pain signals from reaching the brain.</p>	<p>It improves the immune system by raising levels of sex hormones, cortisol, somatostatin, vasoactive intestinal peptide, and cytokine levels.</p>	<p>It stimulates secretion of endorphins (Enkephalins) or regulates receptor activity (mu-opioids receptors) which naturally help with pain.</p>	<p>It affects neurotransmitter levels including serotonin, substance P, dopamine, and Noradrenaline. This would explain the 'qi high'. It also influences immune cells: WBCs, gamma globulins, exomes and mast cell production.</p>	<p>It dilates blood vessels increasing circulation of fresh RBCs and WBCs to an injured area usually by altering calcitonin gene-related peptide (CGRP), prostaglandins, bradykinin, histamine, and angiotensin II</p>

Mechanical Mechanism

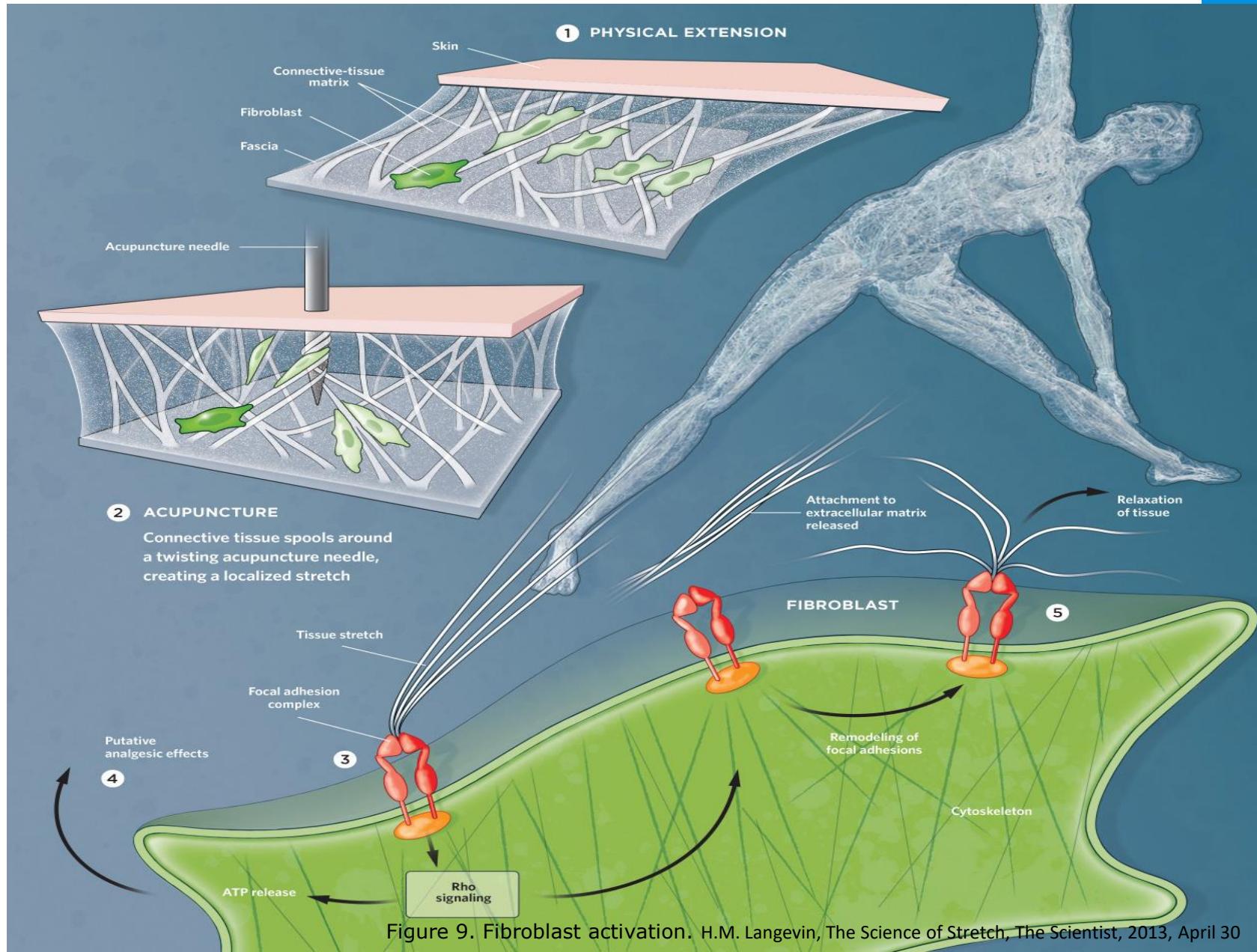
- Langevin & Yandow, 2002, *The Anatomical Record*:
 - Filiform needle presses passes the skin and applies pressure to nerves, connective tissue, fat, etc
 - .16x15mm (diameter: 160 micrometer μm)
 - Whirling action
 - Myofascial Trigger Points
 - Motor Points
 - Up to 2mm – ~10mm insertion
 - Maybe 3-4 inches
 - Acupoint depth vs trigger point/MP

Figure 8. Connective tissue whorl. Langevin & Yandow, 2002.



Mechanical Mechanism

- Langevin, 2013, *The Scientist*:
 - **Pressure & stretch** activate focal adhesion complexes to release **fibroblasts** (mediated by protein **Rho**)
 - Releases **ATP**
 - **ATP** >>> cell shape + analgesia
 - **Rho pathway** remodels adhesions



Mechanical Mechanism

- Langevin, Churchill, Cipolla, 2001, *The FASEB Journal*:
 - **Mechanical load signals** reorganize cytoskeletal tissue.
 - "Intracellular signaling pathways activate **phosphorylation of focal adhesion kinase** and **extracellular regulated kinase (ERK)**."
 - **ERKs** can change gene expression of **collagen XII, tenascin-C, and platelet derived growth factor** via nuclear binding proteins, i.e. NF- κ B
 - I.e. ERKs imp for smooth muscle cell growth and epigenetic changes
 - Rec: seniors need more txs typically

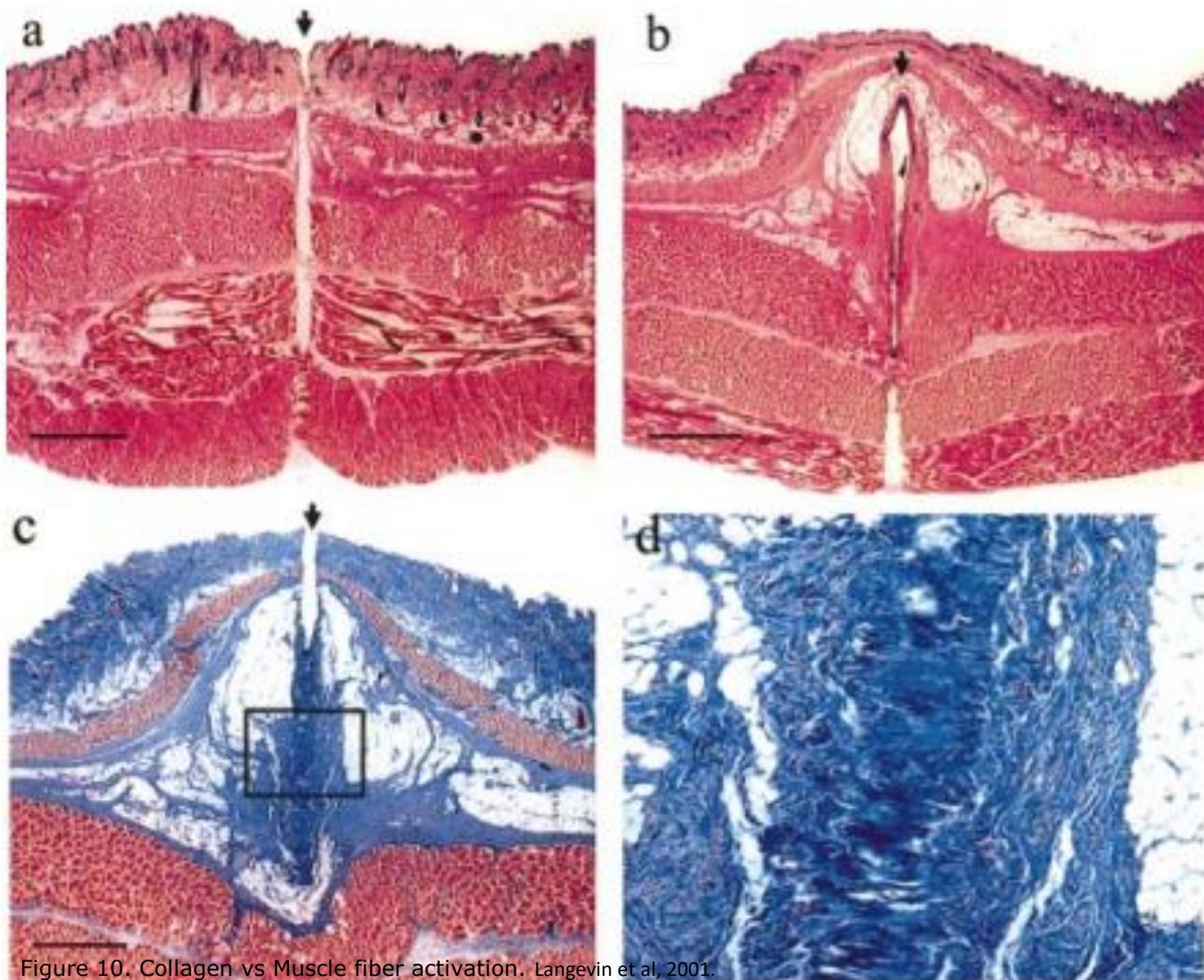
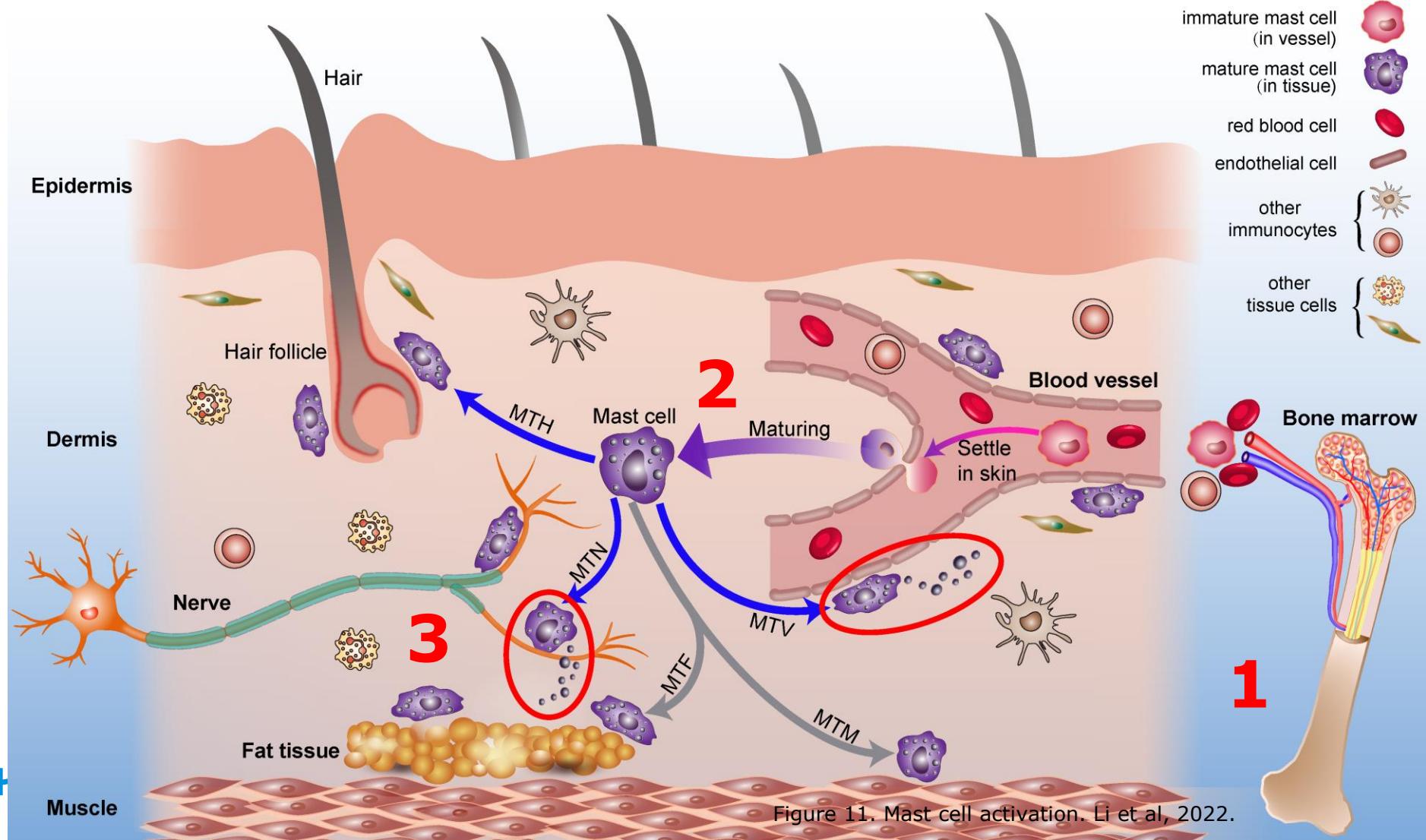


Figure 10. Collagen vs Muscle fiber activation. Langevin et al, 2001.

Mast Cell Mechanism

- Li et al., 2022, *Cells*:
- Local trauma activates **degranulation** of cells which release:
 - Histamine, bradykinin, prostaglandins, CGRP: smooth muscle relaxation via blood vessel dilation
 - Substance P
 - Serotonin
 - B-Tryptase
 - Vasoactive Intestinal Pep
 - Various: IL-1, IL-4, IL-5, IL-13

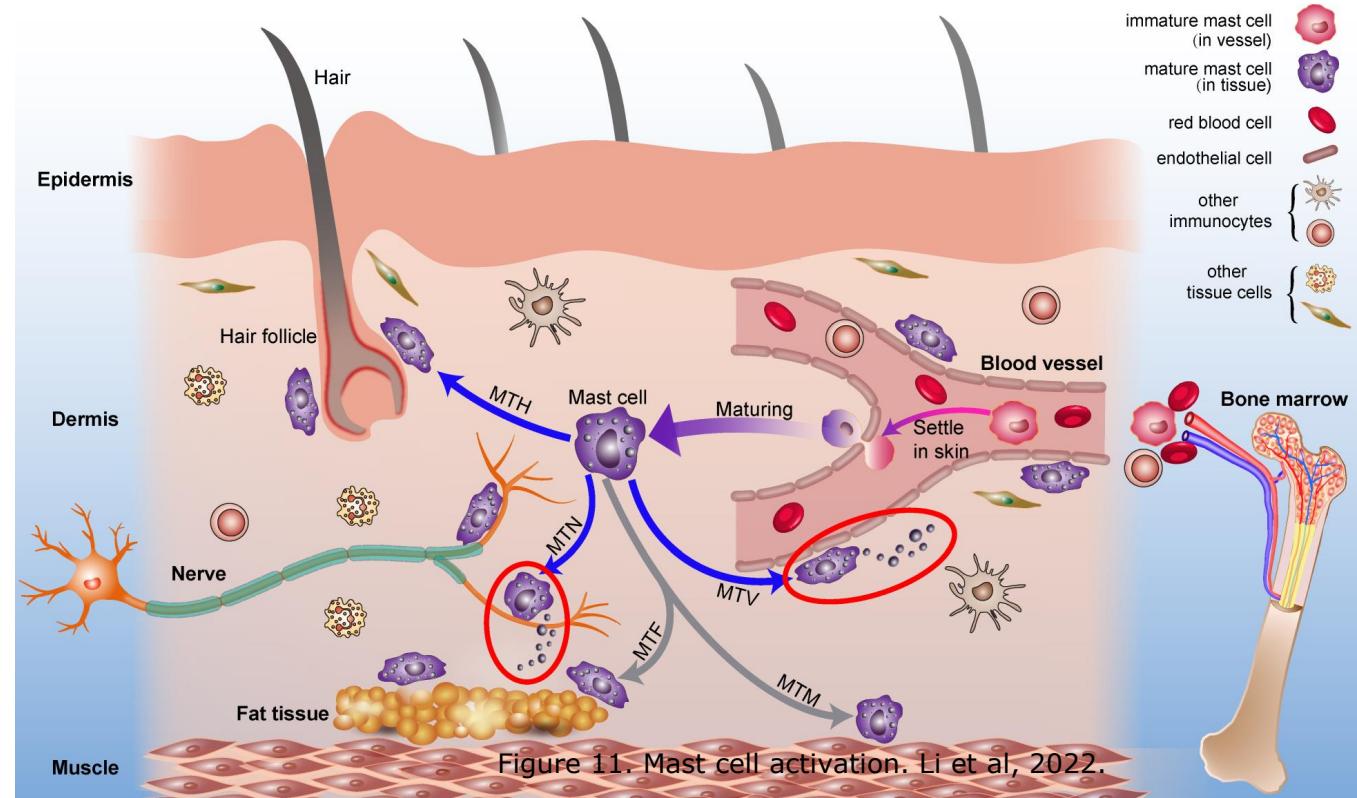
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Mast Cell Mechanism

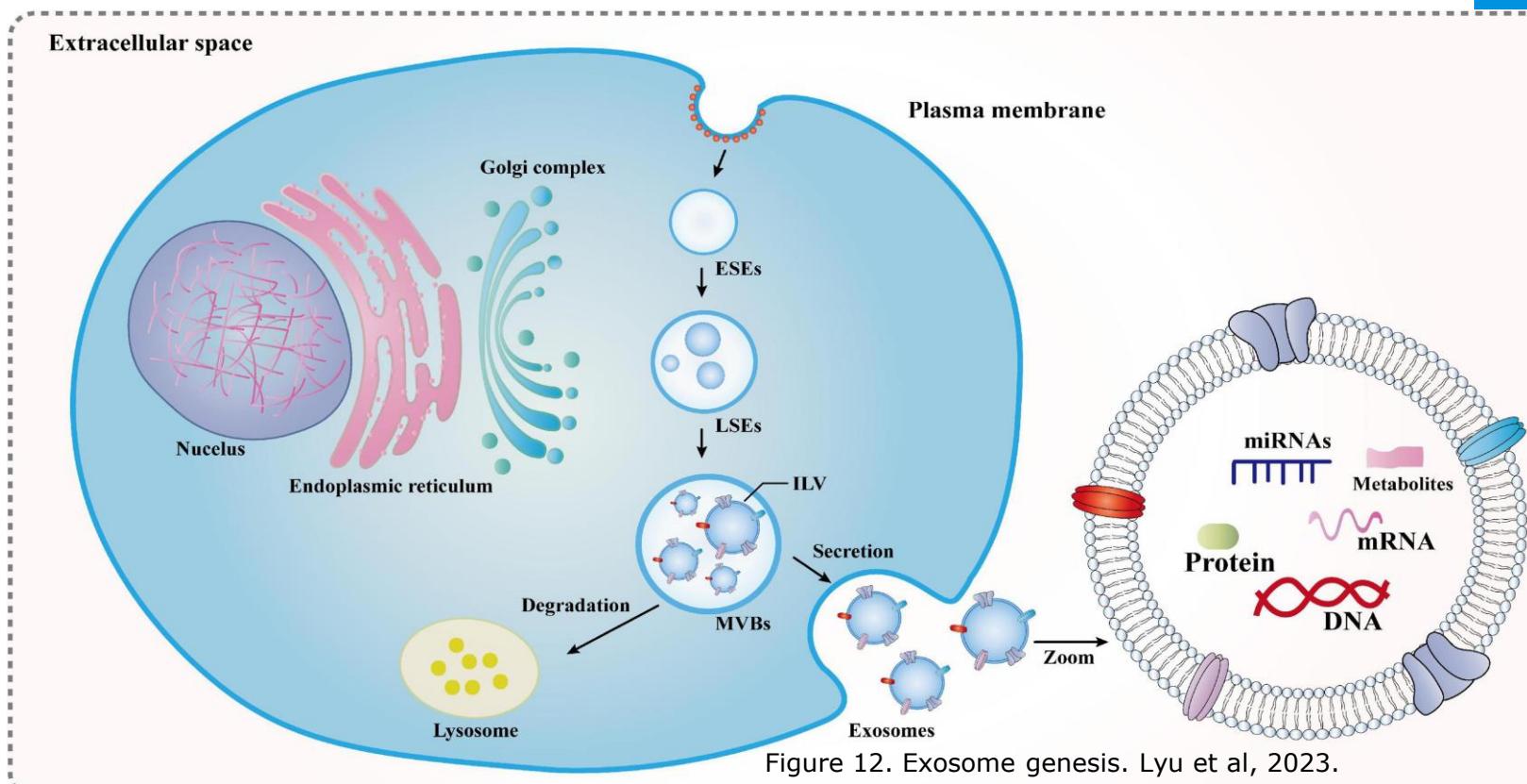
• Li et al., 2022, *Cells*:

- Local damage >>> **degranulation** of mast cells >>> **++ ATP, ADP, AMP, and adenosine** imp in neuropathic and inflammatory pain signaling
- “**ATP** acts as an extracellular messenger to transmit sensory information both at the peripheral site of tissue damage and in the spinal cord.” (Falk, Uldall, Heegaard, 2012)
- **ADP** agonizes **P2Y1 receptor** >>> lowers nociceptive transmitters in spinal cord.
- **Don't drink coffee before your acu treatment!**



Exosomal Mechanism

- Bo et al., 2017, *Chinese Journal of Integrative Medicine*:
 - **Intercellular vesicle messenger** cells released from **mast cells** (also B & T lymphocytes, and **all** cells) that react to stress/invasion >>>> delivers over **400 different kinds of proteins** that regulate our neuro-immune networks.
 - Stimulate sensory nerves to release **Substance P** (neuropeptide) >>> positively reinforces more mast cell activation



Mast Cell Mechanism

- Ming, 2022, *Medical Acupuncture*:
 - Jeimi Song studied mast cell activation, published in 1977 and 1980.
 - First major theory be proven
 - **Degranulation** sig higher in EA and Manual Acu cp to controls
 - Mechanical changes in **collagen fibers** further activate degranulation – positive reinforcing cycle
 - MC map and acu systems:
 - Higher density of MC are in periphery and head
 - **Source points** >>> easy access >>> greatest effect on whole body systems >>> **distal needling**
 - Systemic vasodilation effects: **dysmenorrhea, asthma**, etc

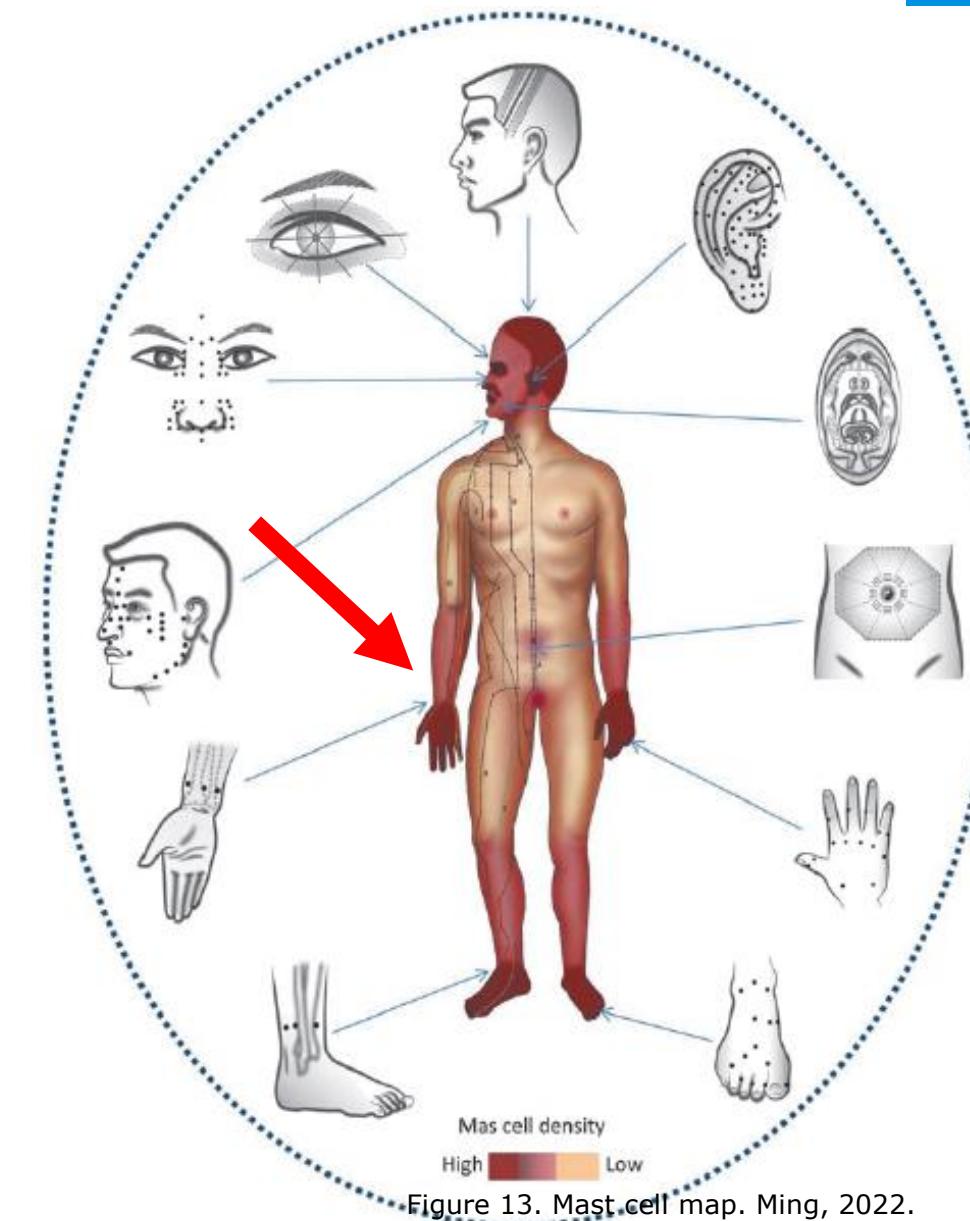
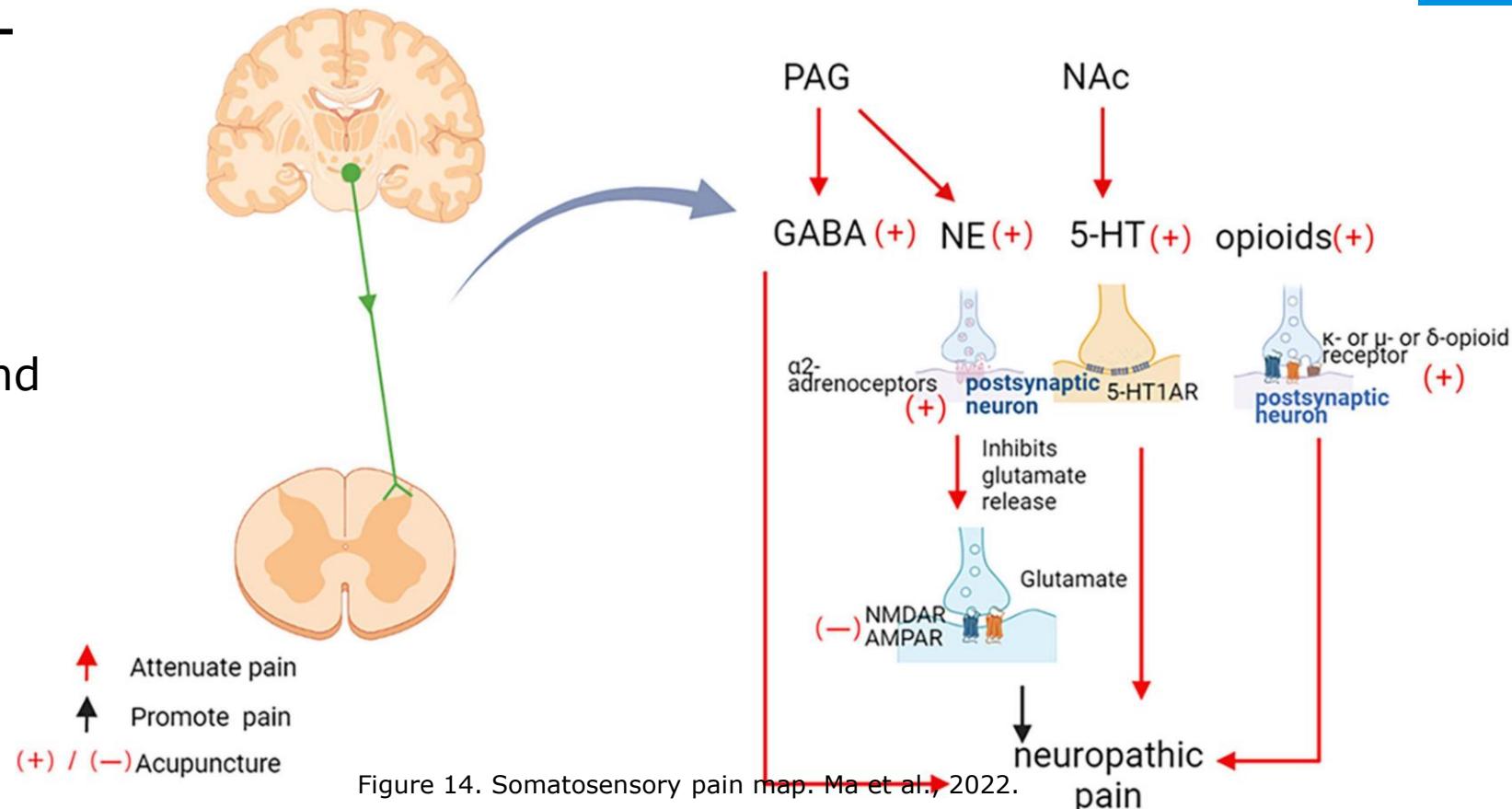


Figure 13. Mast cell map. Ming, 2022.

Neural Mechanism

- Ma et al., 2022, *Frontiers in Neuroscience*, REVIEW, 388 articles:
 - Midbrain and brainstem process nociceptive info via spinal cord
 - **Rostral anterior cingulate cortex** activates **Periaqueductal Gray (PAG)**: Grand central for pain relay
 - **Rostroventral medulla, locus coeruleus** signals release of **GABA, 5-HT, Norepinephrine, and endogenous opioids**
- Opioid peptides enhance gene expression, more opioids express and remain in terminal...
 - More txs >>> greater effect



Neural Opioid Receptor Mechanisms

- Harris et al, 2009, *NeuroImage*:
 - 20 fibromyalgia F patients: **μ-opioid selective radiotracer & PET** (C-carfentanil) to measure MOR sensitivity
 - Found short and long-term modulatory effects of μ-opioid receptor (MOR) binding potential in **amygdala, dorsal and perigenual anterior cingulate**, and **insular cortex**
 - Sham vs Verum: found differences in how μ-opioid receptors became either **MORE** or **less** sensitized
- MacDonald & Chen, 2021, *Frontiers in Neuroscience*:
 - 2 Hz EA associated w/ **met-enkephalin, β-endorphin, and endomorphin** (μ - and δ -opioid receptors) cp. 100 Hz EA stimulates **dynorphin** (κ -opioid receptor)
 - Relationship w/ CB1 and CB2 (cannabinoid) receptors and pain in the **PAG**

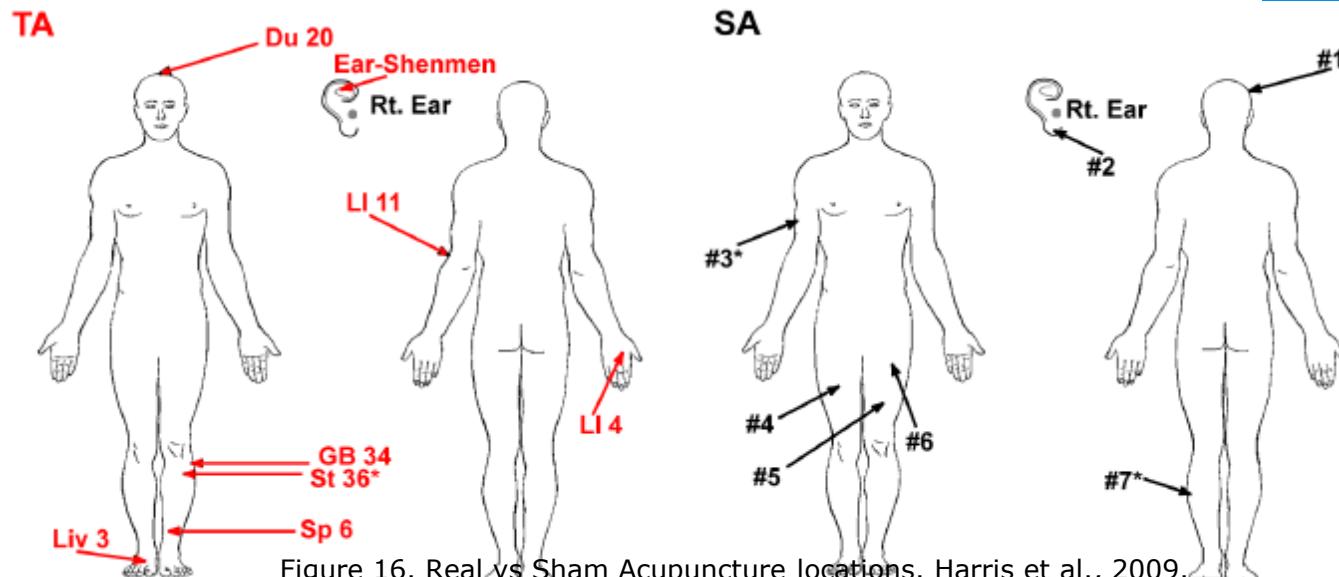
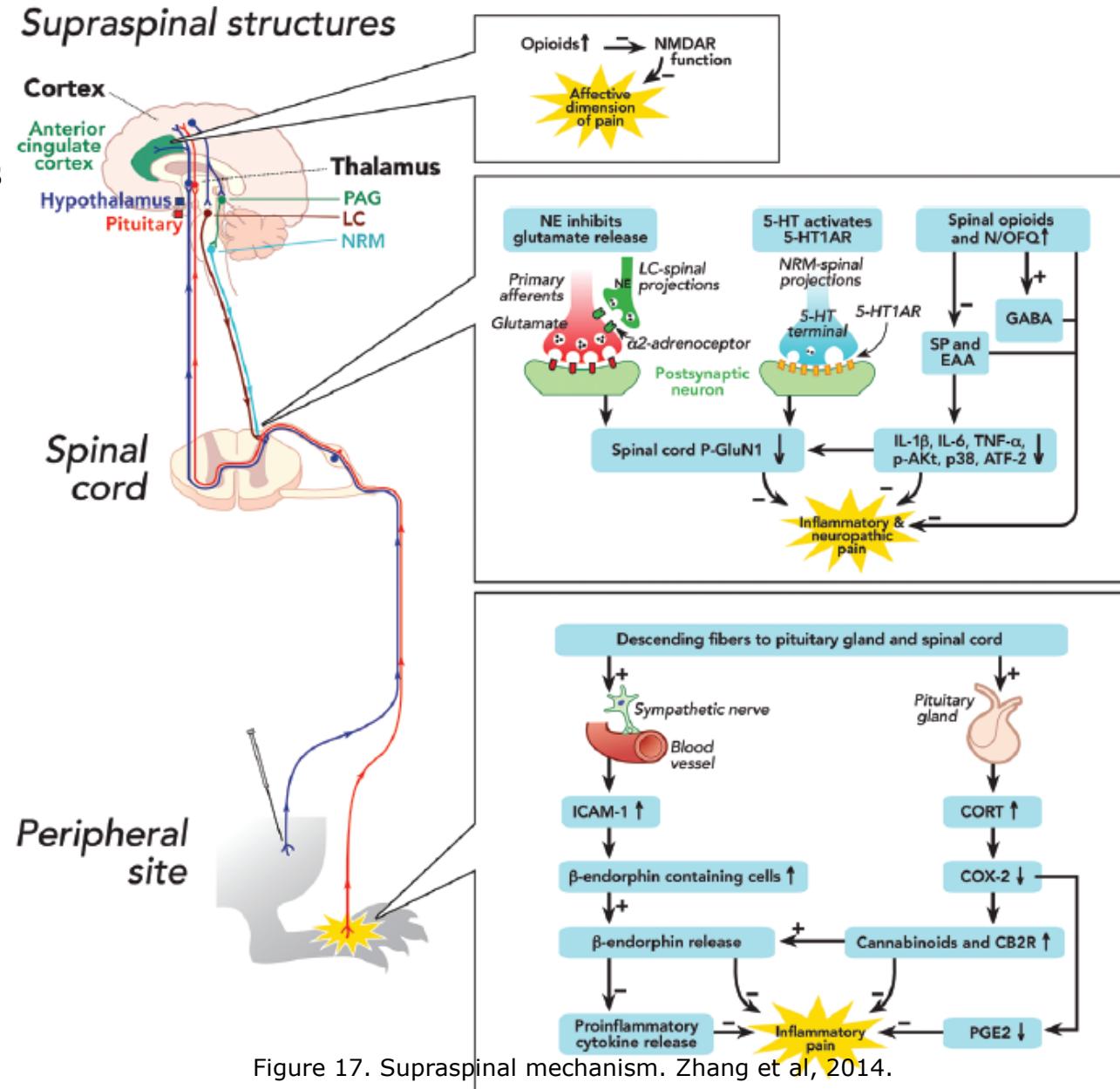


Figure 16. Real vs Sham Acupuncture locations. Harris et al., 2009.

Neural Mechanism

- Zhang et al, 2014, *Anesthesiology*:
 - EA versus manual, 10Hz cp to 100Hz:
 - "...electroacupuncture induces release of **endogenous opioids** from **lymphocytes, monocytes/macrophages, and granulocytes** into inflamed skin."
- Kim et al., 2017: What intrinsic factors influence responsiveness to acupuncture in pain?:
 - High **CCK8** & **naloxone** inhibits effect
 - Some gene and gene expression for proinflam vs anti-inflam: **IL-6, TNF- α**
 - PAG, hypothalamus, spinal cord, pituitary** have central role in mediating pain response



What does it all mean!!!!

- Pressure (**myelinated** A-delta) and pain (**unmyelinated** C) nociceptor neurons activate with needling
 - Local stretch of connective tissue activates **fibroblasts, protein synthesis, ATP, & adaptive epigenetic responses** to reformat local cells
 - At the same time the local tissue damage activates mast cells and **degranulation**, dumping their contents to deal with “invasion.”
 - Neurotransmitters
 - Peptides
 - Cytokines
 - ATP, ADP, AMP, adenosine
 - Histamine, etc, etc, etc
- HENRY FORD HEALTH®**

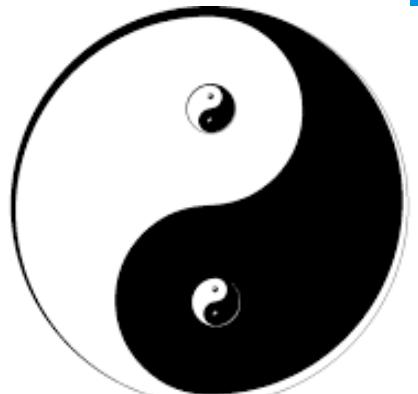
What does it all mean!!!!

- **CNS** responds via **brainstem** and **mid brain** structures, i.e. **PAG**.
- “I’m under attack!”
 - Mounts defense and repair mechanisms: **endogenous opioids, GABA, MOR regulation**, and cortical **functional connectivity (FC)** improves.
- Reduction of inflammatory markers (**IL-6, TNF-α**) and stress hormones (**cortisol**)
- Upregulation of anti-inflammatory **cytokines** like, **IL-10**, etc; and other positive systemic changes in heart rate, BP, HRV.
- EA might have some added benefit

Hormesis

- Calabrese & Mattson, 2017:
 - “Hormesis refers to adaptive responses of biological systems to moderate environmental or self-imposed challenges through which the system improves its functionality and/or tolerance to more severe challenges.”
 - What doesn’t kill you makes you stronger
 - Regenerative medicine
- **Examples:**
 - Exercise
 - Fasting → Yoshinori Ohsumi – 2016 Nobel Prize in Physiology or Medicine
 - Saunas, heat exposure
 - Ice baths, cold exposure
 - Hyperbaric oxygen therapy
 - Cosmetic therapies: i.e. micro dermal abrasion therapy

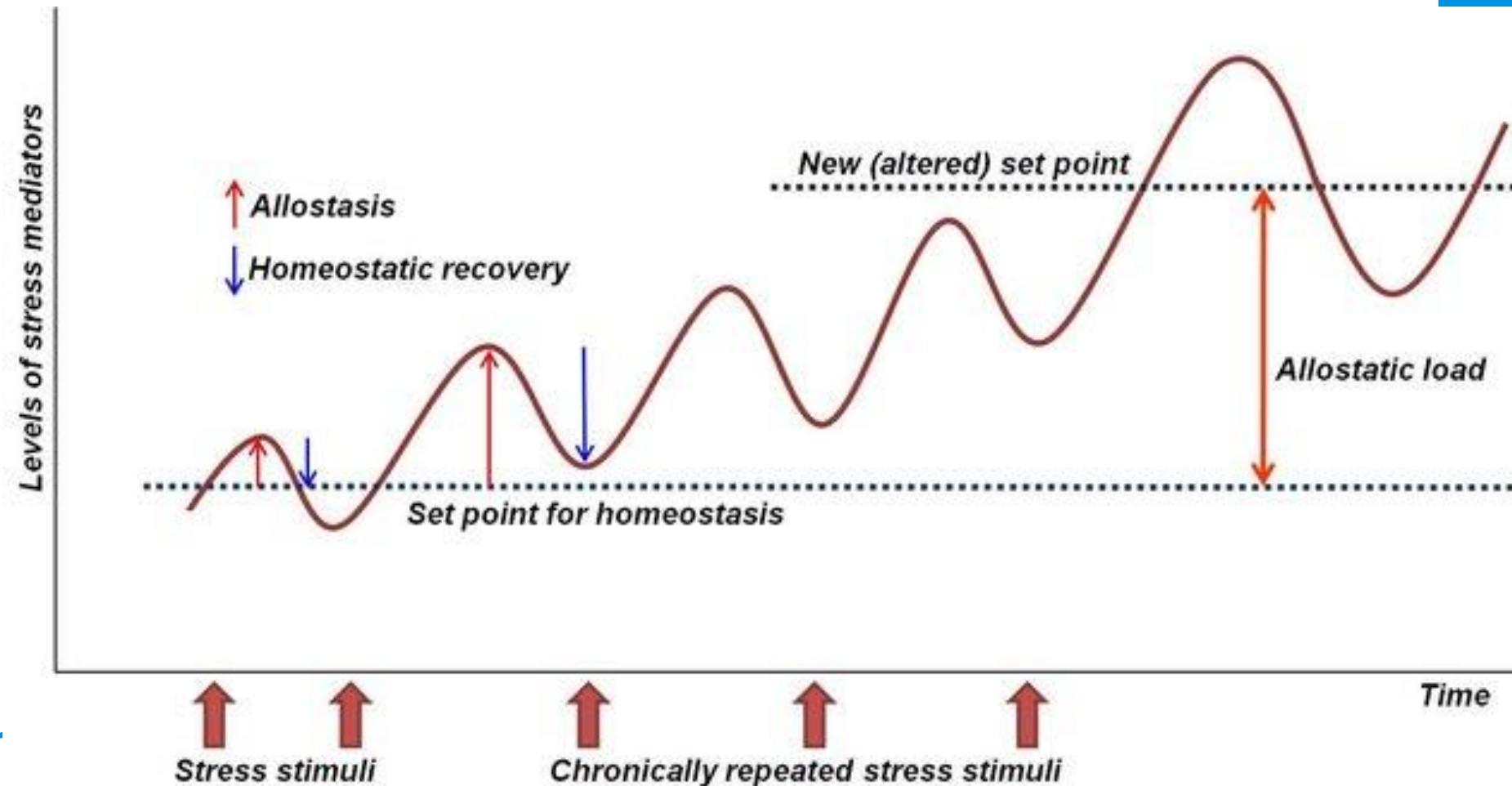
ACUPUNCTURE!!!



Allostasis

- Lee, Kim, & Choi, 2014:
- Chronic stress, pain, disease, weakens ability to return to original homeostatic baseline
- Meanwhile, our ability to handle stress diminishes
- Acu *tricks* the body to produce **slightly** more inflammation to then downregulate
- Training the body to get “there” on its own

Figure 18. Allostasis. Lee, Kim, Choi, 2014.



Pluripotent Mechanisms

Classical



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Norepinephrine

Met-enkephalins

ERKs

Ca²⁺

5-HT

Modern

PAG, rostroventral medulla, locus coeruleus, hypothalamus, spinal cord, pituitary, amygdala, dorsal and perigenual anterior cingulate, insular cortex, etc etc

GABA

β-endorphin

Fibroblasts

Endomorphins

Rho protein

ATP, ADP, AMP

CB1, CB2 receptors

monocytes/macrophages

dynorphin

Norepinephrine

Changes in HRV and heart rate

Nuclear binding proteins: NF-κB

Adenosine

Exosomes

CGRP

μ-opioid receptors

δ-opioid receptors

Mast cell activation

TNF-α

Prostaglandins

Bradykinin

Histamine

Vasoactive Intestinal Peptide

B & T lymphocytes

Substance P

Gene transcription factors: collagen

XII, tenascin-C,

platelet derived growth factor, etc

Interferon-γ

B-Tryptase

Vagal Mechanisms

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Vagal Mechanism

- Adaptive stress responses all processed by the VAGUS
- *Surgically implanted stimulators* (Afra et al., 2021):
 - First device implanted **1988** for epilepsy
 - FDA approval for epilepsy **1997**; depression **2005**
 - FDA approval for gastroparesis in **2000** (Levin, Ortiz, Katz, 2007)
 - FDA approval for stroke affecting up limb (FDA, 2021)

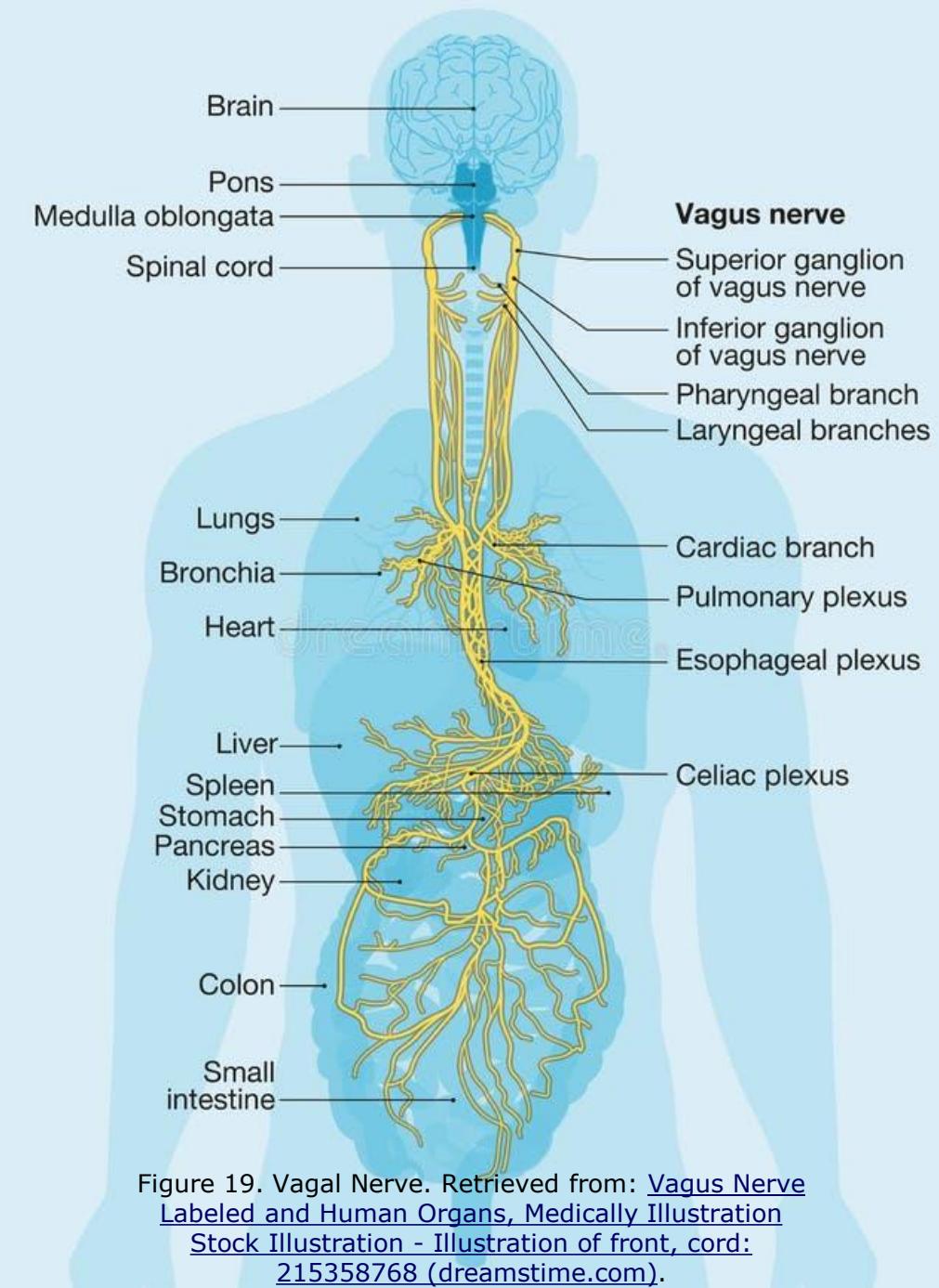


Figure 19. Vagal Nerve. Retrieved from: [Vagus Nerve Labeled and Human Organs, Medically Illustration Stock Illustration - Illustration of front, cord: 215358768 \(dreamstime.com\)](https://www.dreamstime.com/vagus-nerve-labeled-and-human-organs-medically-illustration-stock-illustration-illustration-front-cord-215358768-dreamstime.com).

Vagal Mechanism

- Auricular Branch of the Vagus Nerve (ABVN):
 - **Spinal trigeminal nuclei of the medulla** and **nucleus of the solitary tract (NTS)**, provides regulation of **locus coeruleus (LC)** (Vargas-Caballero et al., 2022)
 - **LC**: Adaptation of stress, panic via noradrenaline regulation
- **Transcutaneous auricular** vagal stimulation:
 - Pain modulation (Greif et al., 2002)
 - Obesity (Clancy et al., 2014)
 - Migraines (Luo et al., 2020)
 - Depression (Wu et al., 2018)
 - Anxiety (Usichenko et al., 2022)
 - Insomnia (Jiao et al., 2020)
 - POTS (Diedrich et al, 2021)



Figure 20. Auricular Branch of the Vagus Nerve. Clancy et al., 2014.

Vagal Mechanism

- *Transcutaneous* vagal stimulation:
 - Alzheimer's (Vargas-Caballero et al., 2022)
 - Lowered IL-1, IL-6, TNF-a; increased: BDNF
 - Sepsis (Torres-Rosas et al., 2014)
 - Lowered TNF-a, IL-6, MCP-1 and INF- γ
- Cervical stim:
 - **PTSD** (Bremner et al., 2020)
 - Blocked IL-6 and IFN- γ from rising in stressful situations
 - Do before schedule 1 drugs?
 - **TBI** (Zhou et al., 2014)
 - TNF-a, interleukin-1 β (both fell) & IL-10 (rose)
 - Both might have neuroprotective effects



Figure 20. Auricular Branch of the Vagus Nerve. Clancy et al., 2014.

Vagal Mechanism

- Peng et al., 2017: Tinnitus

AcuPUNCTURE + vagal stimulation

- Napadow et al, 2012:

- Endometrial pelvic pain patients, n = 15
- ACU + VAGUS STIM w/ exhalation phase of breathing versus non-vagal stimulation
- Pelvic pain decreased independent of anxiety decreasing
- Measured: respiration, ECG (QST), HRV, and heart rate

- Case study: Hyperthyroid patient

- 30Hz cp to ~5Hz

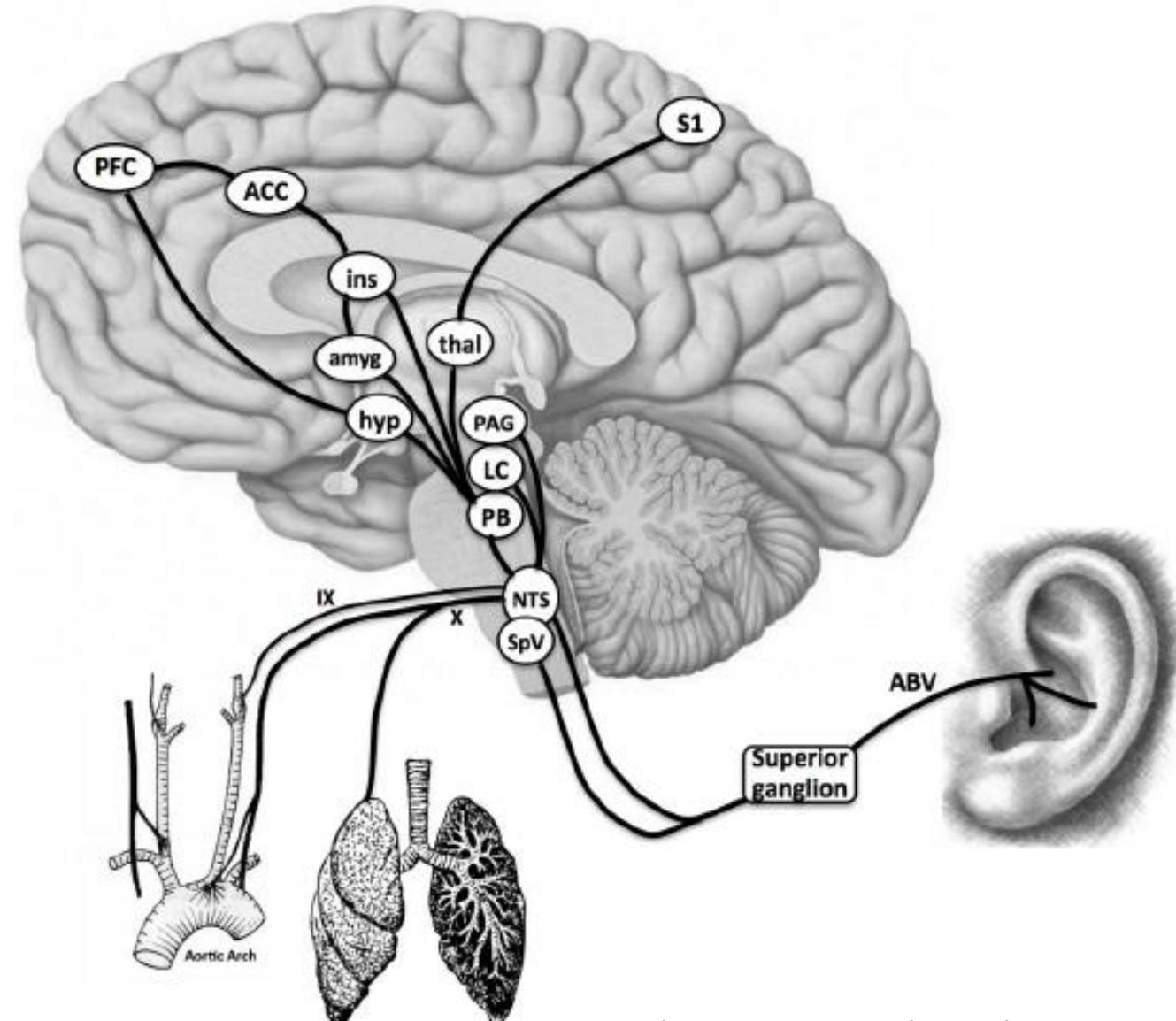


Figure 21. Nucleus tractus solitarius image. Napadow et al., 2012.

Vagal Mechanism

AcuPUNCTURE + vagal stimulation

- Boehmer et al, 2020: found modest impact on HRV
- Winkler et al, 2021: somatic cell count lowered in cow mastitis
 - Love non-human studies, no placebo!

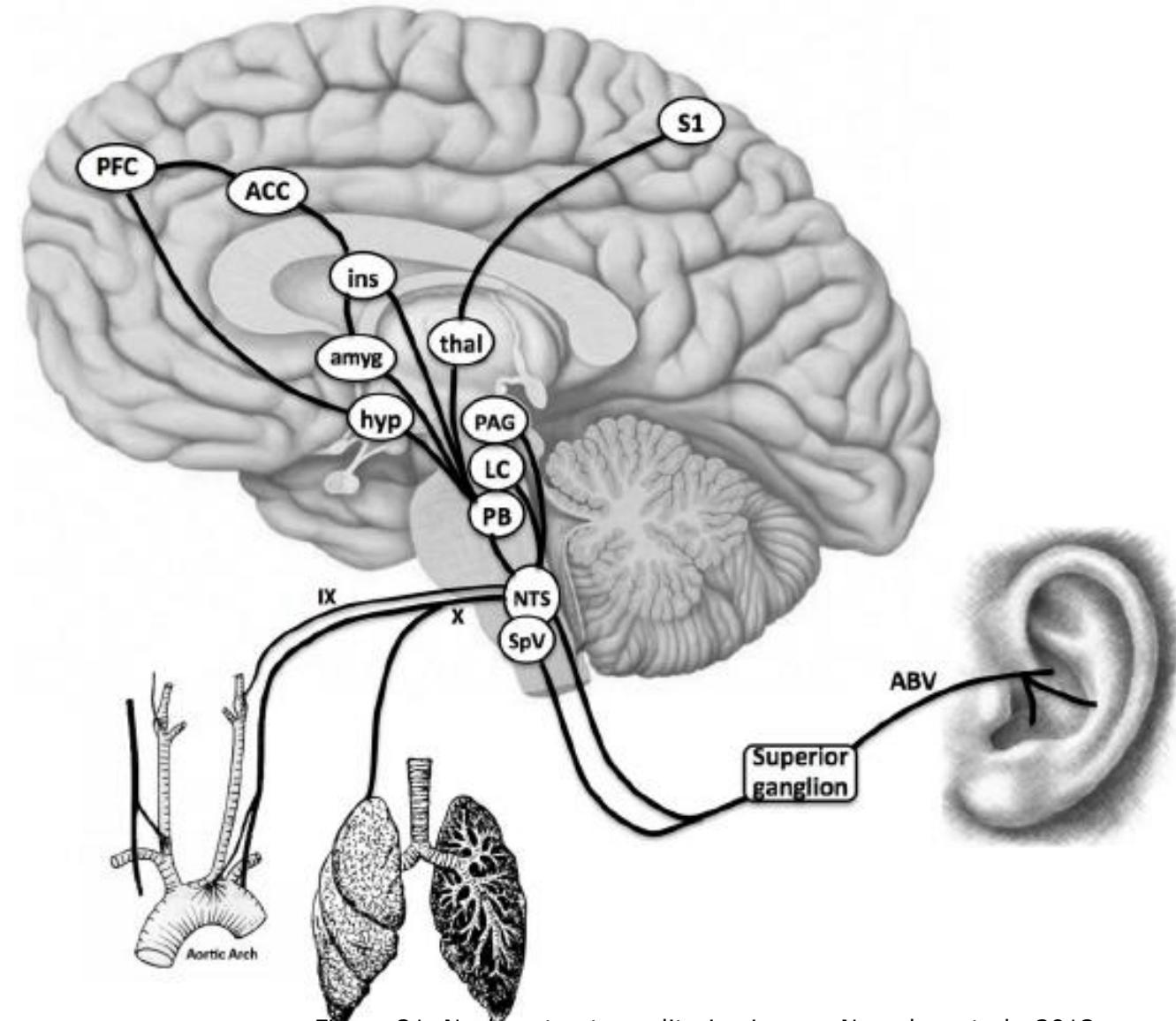


Figure 21. Nucleus tractus solitarius image. Napadow et al., 2012.

But wait! There's more!

Vagal Mechanism

- Liu et al, 2021 in *Nature*
 - EA at ST36 lowered TNF and IL-6 via vagal adrenal axis, but not in **genetically ablated PROKR2Cre-gene** marked sensory neurons, in mice
 - Cp to ST25, an abdominal point, sympathetic reflex was activated
 - Demonstrates specificity of reactions at different regions
- Notable mention:
 - Du & Liu, 2015: EA at ST36 induced enteric neuro regeneration, rat model

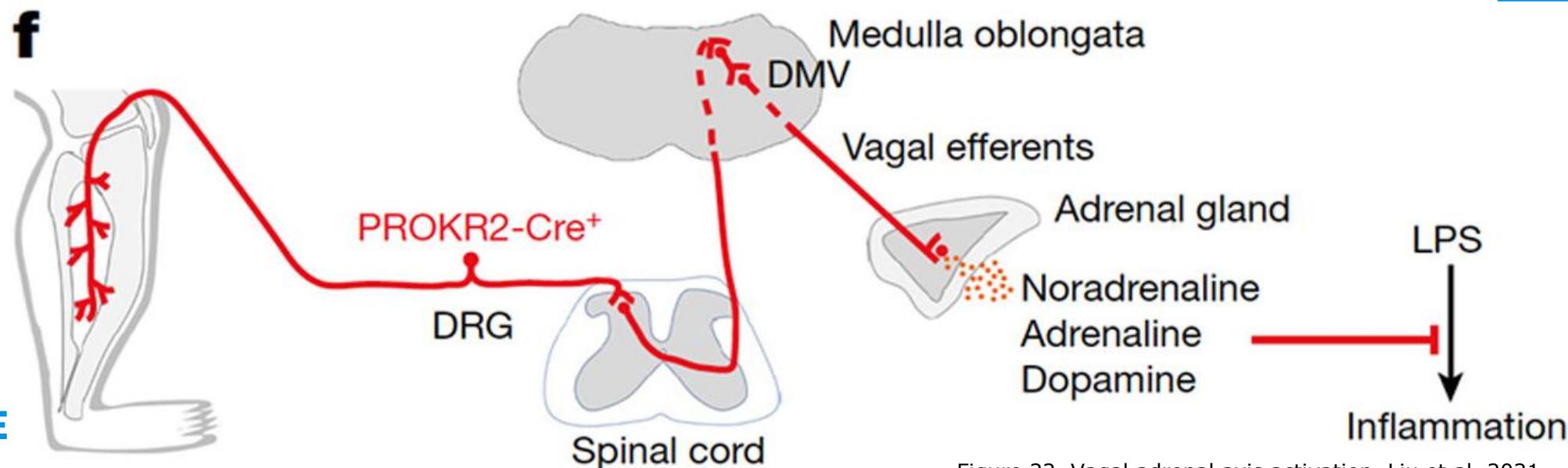


Figure 22. Vagal adrenal axis activation. Liu et al, 2021.

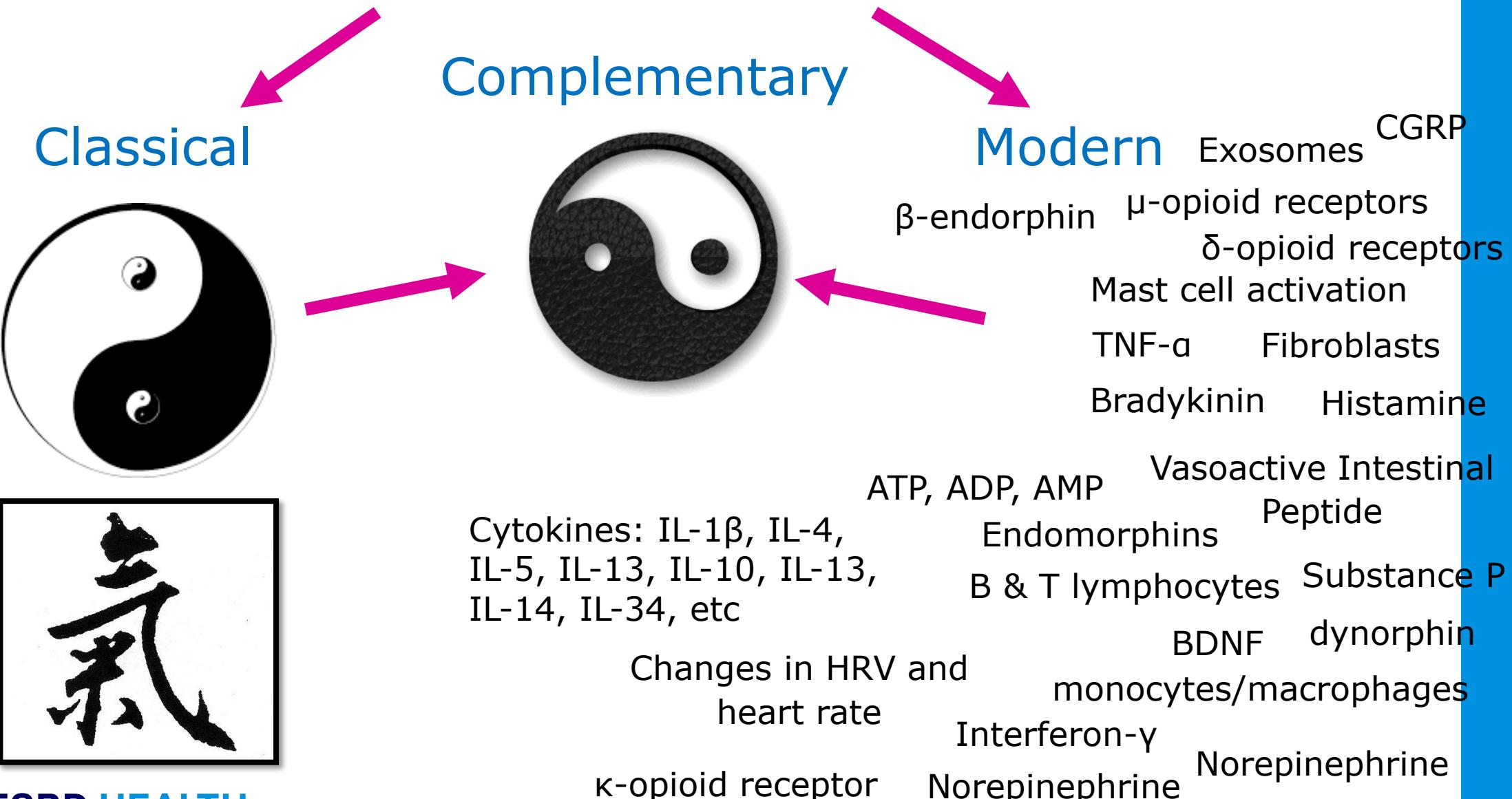
Why is this important?

- Can the body heal when in chronic fight/flight/freeze?
 - Full body acupuncture
 - Auricular acupuncture
 - Transcutaneous stimulation
 - Breathing exercises, i.e. qi gong, martial arts
- Body remembers emotions and higher states of being

Integrative Care & Other Translations

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Comfortable with Translations



Dry Needling Translations

- Legge, 2014:
 - Dr. Janet Travell published in 1941
 - Brav & Sigmund's 1941 paper references acupuncture's role in low back and sciatic issues
- SI12 over supraspinatus MP
 - Indications, intention
- Quiz time:
 - A. Stuck qi in Small Intestine
 - B. Tendinopathy in the supraspinatus

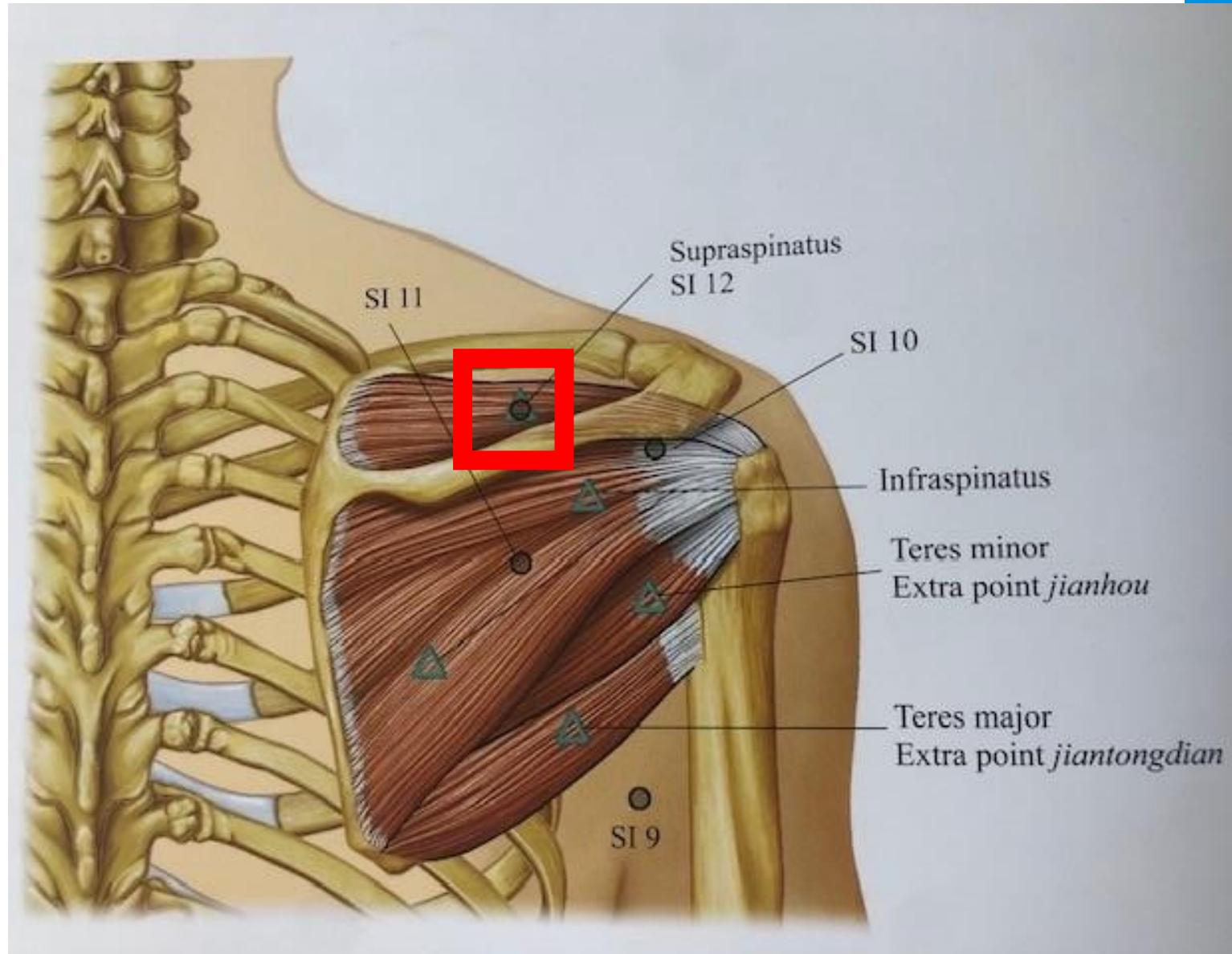
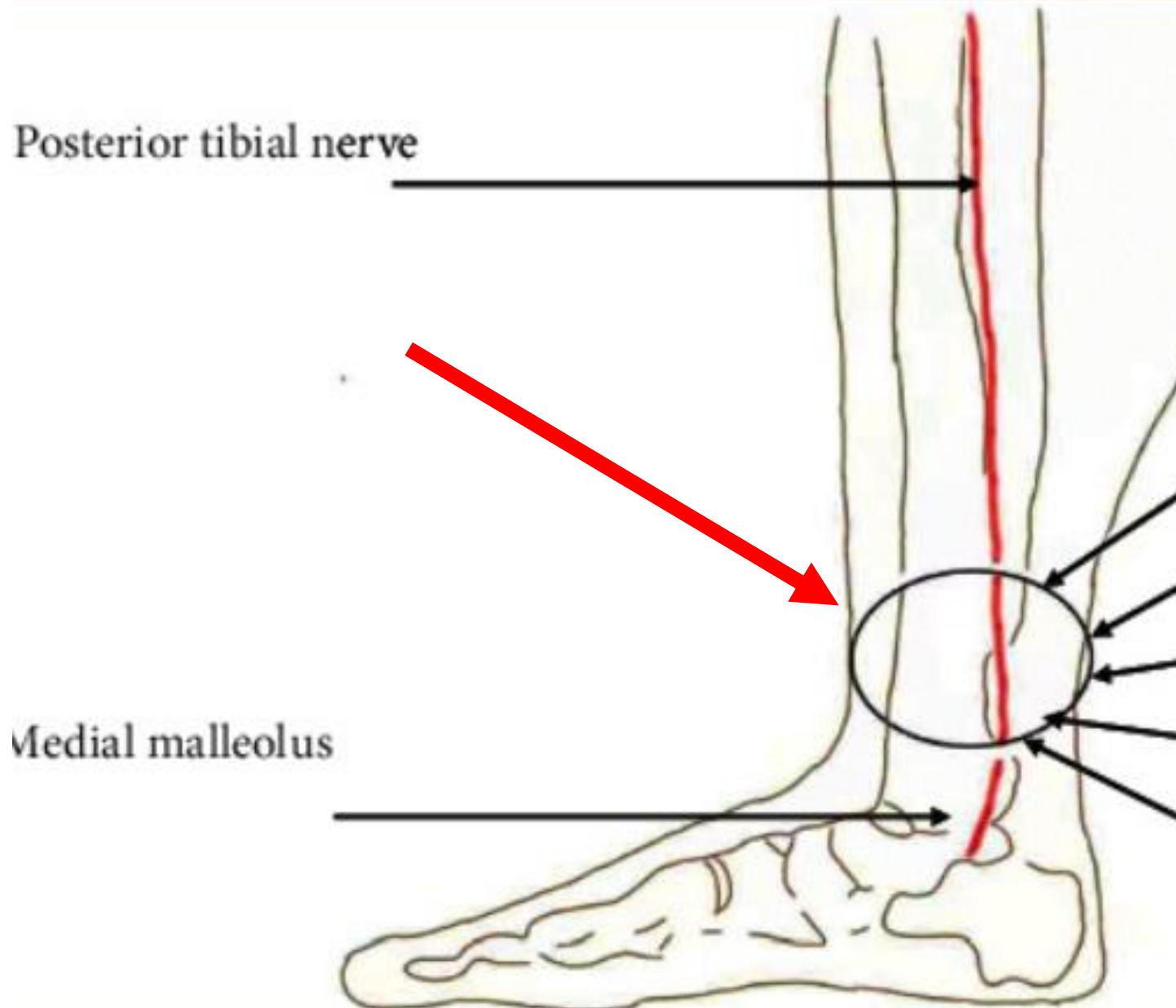


Figure 23. Scapula muscles. Callison, 2019.

Translations

Figure 25. PTNS. Al-Danakh et al., 2022.

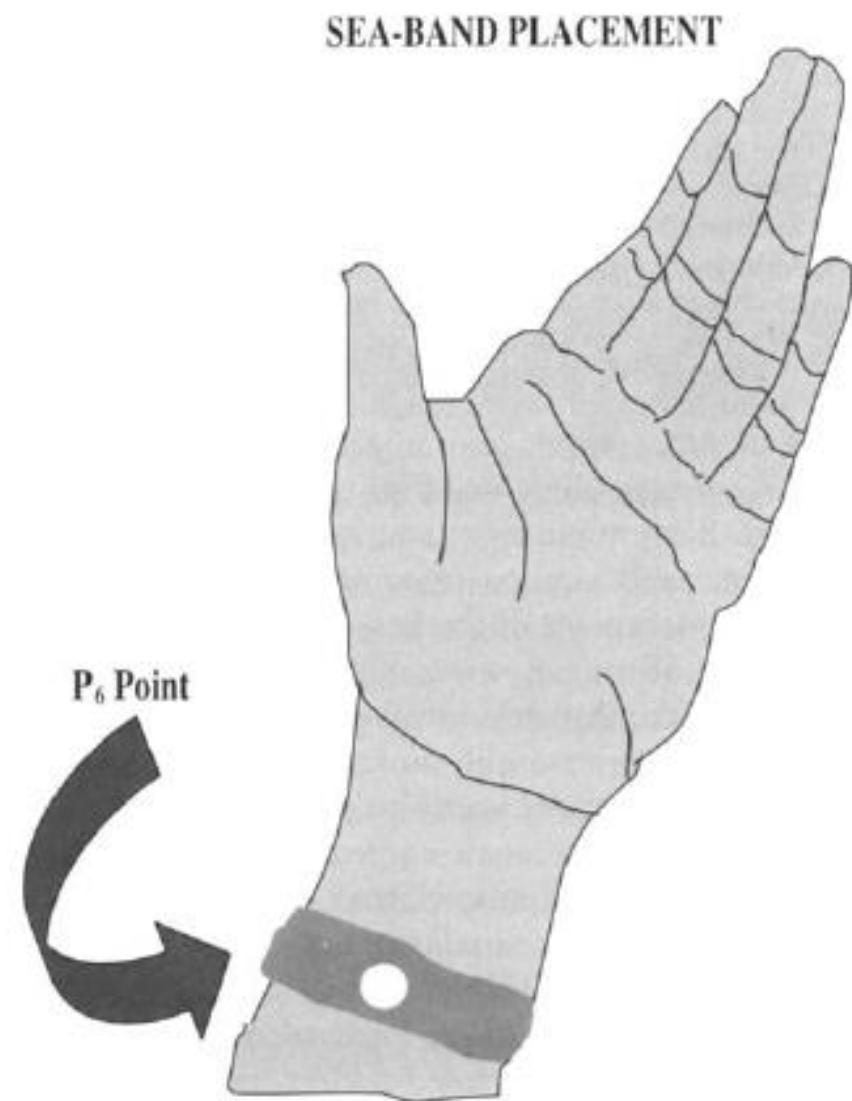
- Posterior Tibial Nerve Stimulation (PTNS) for Over Active Bladder (OAB):
 - Dr. McGuire developed from acupuncture techniques in **1983**
 - L4-S3 nerve root pathway:
 - PTN
 - Pelvic floor
 - Bladder sphincters
 - Bladder detrusor muscle
 - (Al-Danakh et al., 2022)
 - CMS reimbursable



Translations

- Sea Bands:
 - Pericardium 6 >>> pressure on median nerve
 - **Motion sickness**, nausea,
pregnancy nausea/vomiting, post-operative nausea, CT nausea
- Steele et al, **2001**, *Journal of Obstetric, Gynecologic, & Neonatal Nursing*

Figure 26. Sea-band placement. Steele et al, 2001.



Translations

- ReliefBand, “FDA-cleared”:
 - Coloma et al., **2002**, *Anesthesiology*:
 - Acustim + Zofran = enhanced efficacy
40% vs 73%
- Bai et al., 2010, *Journal of Magnetic Resonance Imaging*:
 - Evidence of activation: “cerebellar-hypothalamus and insula”
 - Vestibular receptors and proprioceptors

Figure 27. *Reliefband device product lineup*.
Retrieved from [Reliefband®](#)

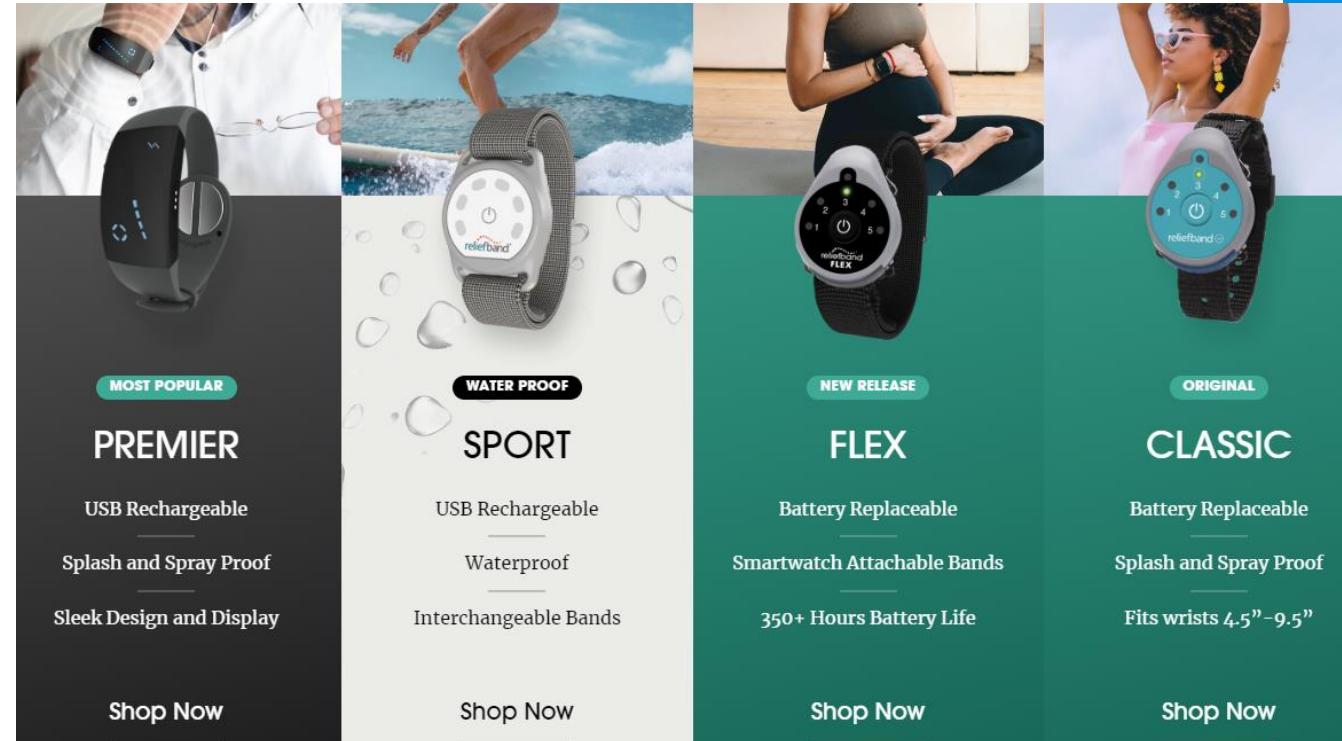
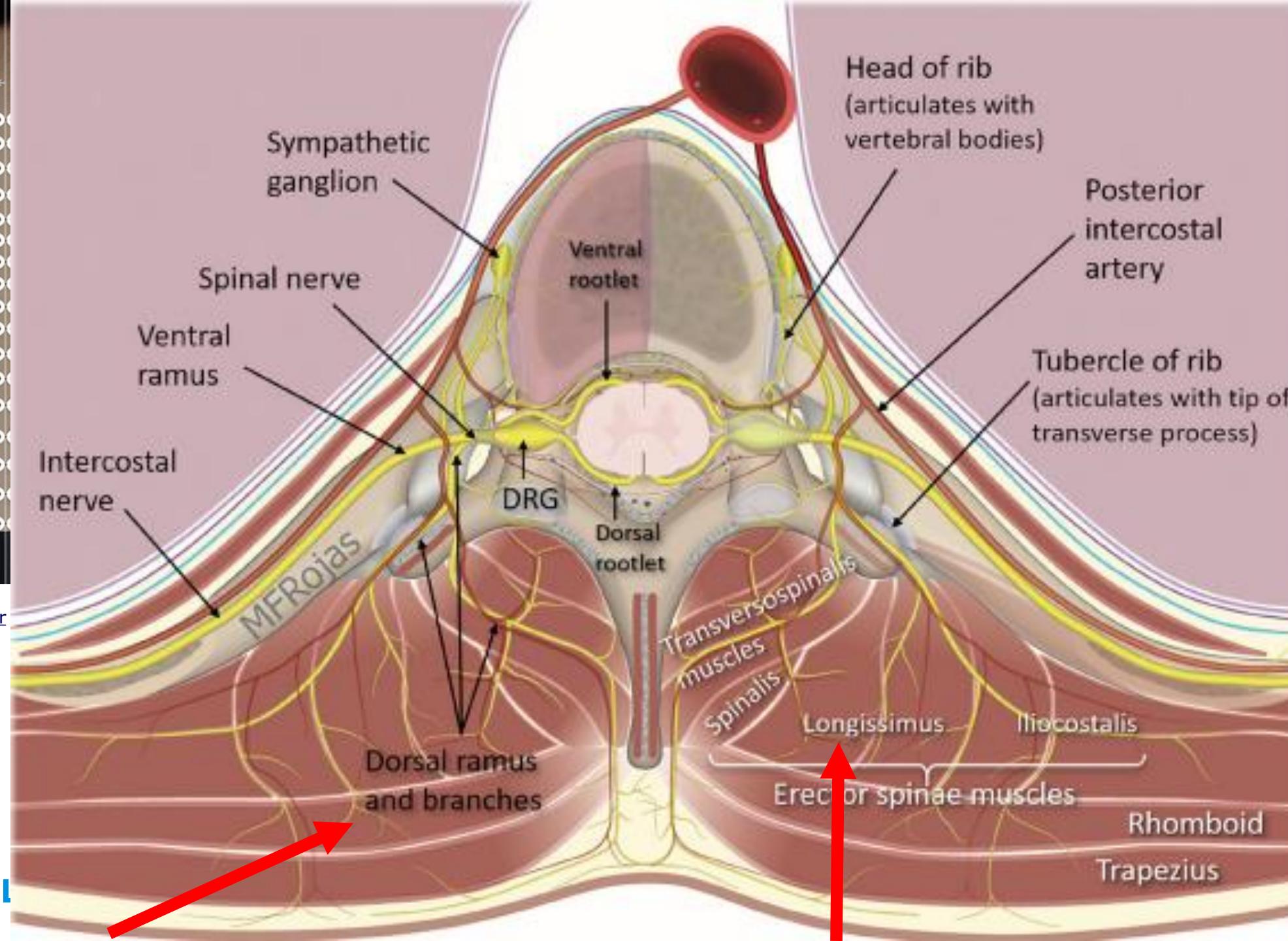




Figure 28. Hua tou jia ji point location.
Retrieved from [Huatujiaji \(Ex-B2\) | Master Tung's Acupuncture | eLotus CORE \(mastertungacupuncture.org\)](#).



More on acupuncture informing biomedicine...

- Society of Acupuncture Research's (SAR) White Paper, (MacPherson et al., 2016):
 - Endorphins and endogenous opioids
 - Connective tissue research
 - Placebo response:
 - Provider-patient relationship
 - ID'd biomarkers related to placebo
 - Animal studies
 - Acupuncture devices:**
 - TENS was developed by Dr. C. Norman Shealy, MD in the 1970s
 - Later created non-penetrating devices

Looking towards future understanding

- **National Center for Complementary and Integrative Health (NCCIH)** lead by Dr. Helene Langevin
- Topological Atlas and Repository for Acupoint research (TARA):
 - “1) an acupoint ontology using both Traditional East Asian Medicine (TEAM) and conventional biological nomenclature systems,
 - 2) male and female standardized human and rat body atlases with 3D coordinate system, and
 - 3) an easily searchable database, curated by an expert committee, that contains previously published physiological data associated with acupoint stimulation.”
- (National Institutes of Health, n.d.)
- Important for future Acu research...

Summary and Safety

HENRY FORD HEALTH[®]

QUIZ!

- A. Mechanical
- B. Neuro
- C. Immune
- D. Vagal, restore parasympathetic dominance
- E. All the above
- F. All that we don't yet know

Key Takeaways

- We HAVE evidence to show effect!
- Difficult to translate when speaking different languages
- Hormetic, adaptive, regenerative, salutogenic
- Durability of symptom relief if systemic changes happen
- Don't wait to refer when traditional therapies are refractory

Patient Responsiveness

- Various:
 - Age, general health and established lifestyle
 - Complexity of history
 - Medications: Hurdle OR Synergistically
 - Might not be the right therapy for the patient at this given time
 - MS and erythromelalgia px
 - Don't let this stop you from considering acu, we never know how someone responds until after a few treatments

Safety

- Adverse events are **low** (Chan et al., 2017, *Scientific Reports, Nature*)
 - Auricular Vagal
 - Could benefit heart failure pts: Boosts Nitric Oxide & enhanced peripheral perfusion (Kaniusas et al., 2019)
- Cautions:
 - Pregnancy, certain points avoided
 - Post surgical timeframe varies
- Contraindications:
 - Pacemaker or implanted stimulators, will avoid EA but manual acu is fine

Pricing & Insurance

- Cash rates: \$139 initial
 - \$88 for follow up / \$440 for 6 visits
- Smoking package: 5 for \$200
- Insurances:
 - Private health plans might have benefits, in- and out- of network
 - Superbills
 - VA, WC, MVA
 - Medicare

Learn More

- National Center for Complementary and Integrative Health
 - www.nccih.nih.gov
- Acupuncture Now Foundation
 - www.acunow.org
- Acupuncture Evidence Project, full report:
 - https://acupuncture.org.au/publicassets/1c94b136-802d-ed11-9115-00505687f2af/Acupuncture-Evidence-Project_Mcdonald-and-Janz -Revised-Edition-07-02-17-Referenced.pdf
- *Neuropuncture: A Clinical Handbook of Neuroscience Acupuncture*
 - By Michael Corradino
- *The Web That Has No Weaver* by Ted Kaptchuck

Thank you

mkulas1@hfhs.org

HENRY FORD HEALTH®

- Sit up straight
- Eat your vegetables
- Get enough sleep and water
- Call a loved one

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