WARNING

This material has been reproduced and communicated to you by or on behalf of the University of Melbourne in accordance with section 113P of the *Copyright Act 1968* (Act).

The material in this communication may be subject to copyright under the Act.

Any further reproduction or communication of this material by you may be the subject of copyright protection under the Act.

Do not remove this notice

Veterinary Bioscience: Cells to Systems

VETS30029 / VETS90121













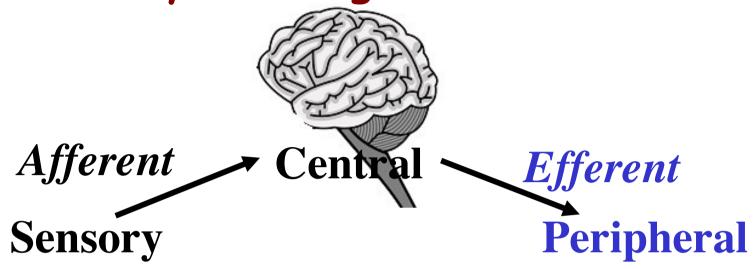
Introduction to the peripheral nervous system – autonomic and somatic nervous systems

A/Prof. James Ziogas
E: jamesz@unimelb.edu.au

At the end of this lecture, you should be able to:

- Describe the basic anatomy of the Peripheral NS (efferent), the Autonomic nervous System (ANS) and its Parasympathetic and Sympathetic divisions
- Describe the major neurotransmitters and receptors involved in chemical transmission within the ANS and somatic NS
- List examples of agonists and antagonists that have selective activity at adrenoceptors

Nervous system organisation





Somatic/Motor (Voluntary)

Autonomic
(sympathetic & parasympathetic)
(Involuntary)

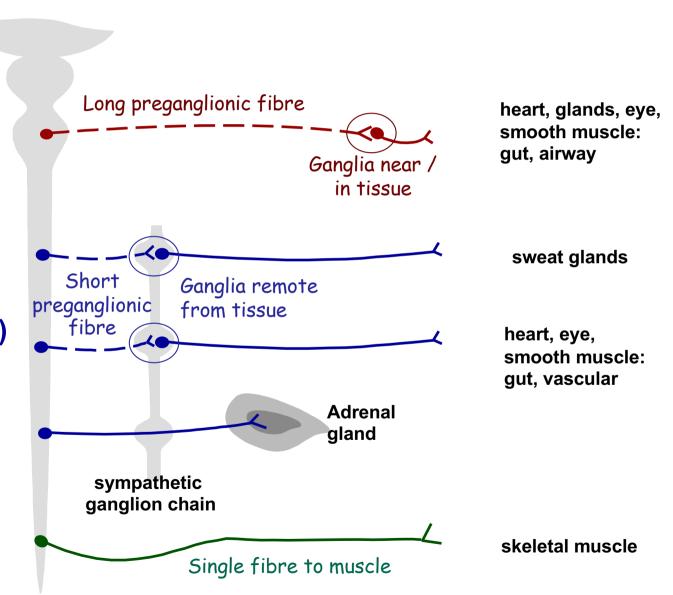
Peripheral Nervous System - Anatomy

Autonomic

Parasympathetic (Cranio/Sacral)

Sympathetic (Thoracic/Lumbar)

Somatic Motor



Peripheral Nervous System - Physiology

Somatic - skeletal muscle → posture & movement

Autonomic - ALL other tissues → homeostasis (housekeeping)

Parasympathetic



- anabolic, rest and repose

Sympathetic

- catabolic, fight or flight
 - neural & humoral elements



Postganglionic Tissue Responses in the Autonomic Nervous System

Parasympathetic Sympathetic (catabolic actions) (anabolic actions)

Heart rate Trate

Arteries constriction

dilation

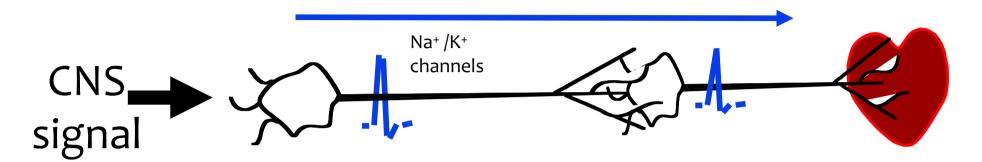
Bronchi constriction dilation

GIT **†** activity **↓**activity

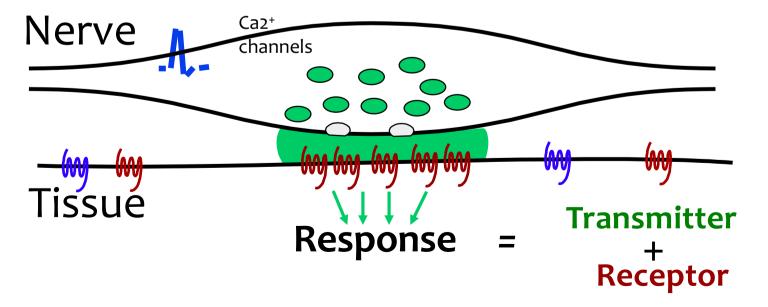
Pupil constriction dilation

Electrical and Chemical signalling in nerves

Action Potential - electrical along nerve



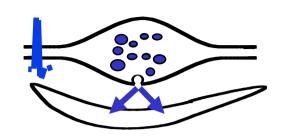
Neurotransmitter - chemical at nerve ending



Chemical transmission in peripheral nerves

Target cell

- Neurotransmitters must bind
 - receptors on target



Learn

- Muscarinic, nicotinic
- α & β adrenoceptors

Signalling cell (nerve)

- · Neurotransmitters need to be:
 - present
 - Synthesised / stored
 - released
 - inactivated

Learn

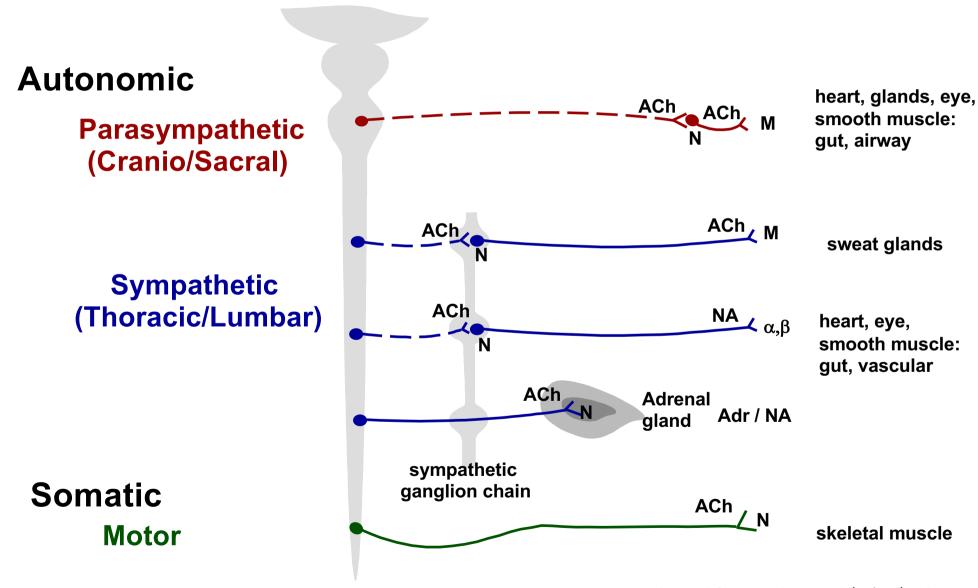
- Acetylcholine
- Noradrenaline
- Adrenaline

At the end of this lecture, you should be able to:

- Describe the basic anatomy of the Peripheral NS (efferent), the Autonomic nervous System (ANS) and its Parasympathetic and Sympathetic divisions
- Describe the major neurotransmitters and receptors involved in chemical transmission within the ANS and somatic NS
- List examples of agonists and antagonists that have selective activity at adrenoceptors

Peripheral Nervous System

- Anatomy, Physiology & Pharmacology



Adapted from Golan et al (3rd Ed); Ch 8

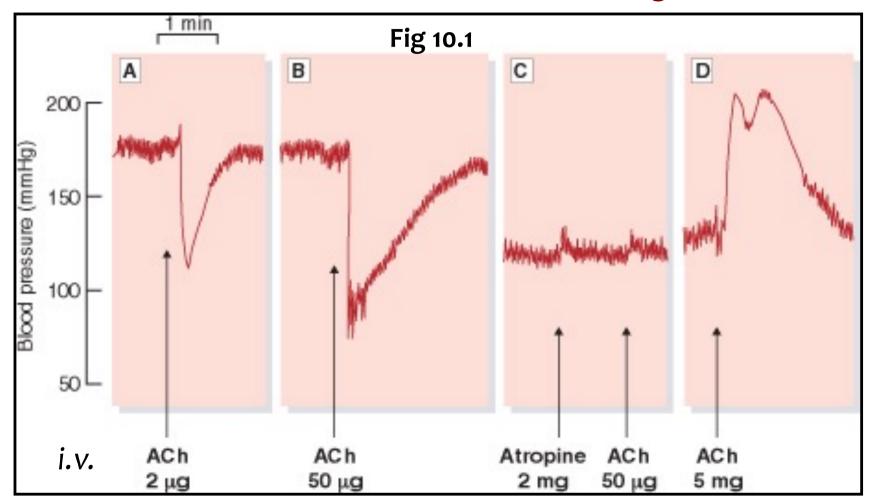
Postganglionic Tissue Responses in the Autonomic Nervous System

Parasympathetic Sympathetic (catabolic actions) (anabolic actions) **T**rate Heart rate **Arteries** constriction dilation (Adr) dilation (Adr) constriction Bronchi **1** activity **GIT L**activity dilation constriction Pupil acetylcholine noradrenaline Transmitter muscarinic Receptor α or β

Effects of Ach On Cat Blood Pressure

A & B Dose-response

C & D Antagonist selectivity



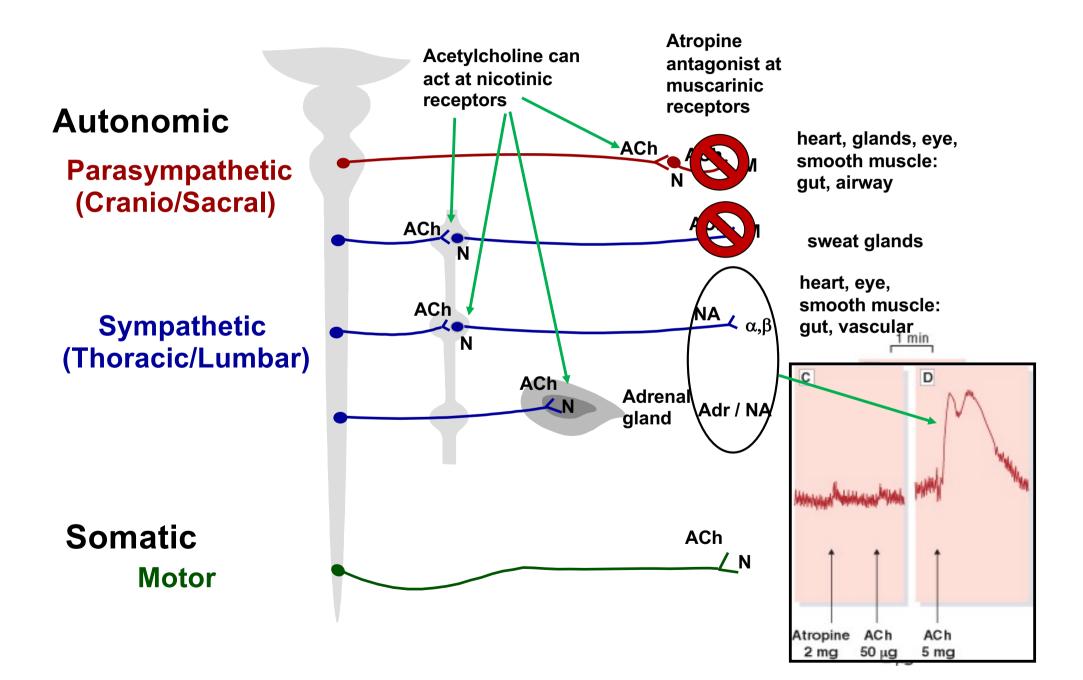
Rang et al; Pharmacology 6^h Edition; Churchill Livingstone, 2007

Receptors: Tissue distribution and innervation

Muscarinic receptors on endothelial cells release NO

Drugs: Action & selectivity

Peripheral Nervous System Anatomy, Physiology, Pharmacology



Cholinergic receptor localisation and action

Autonomic Nervous System

Muscarinic receptors

- Parasympathetic neuroeffector junction
 - Slow responses G protein-coupled, second messengers
 Salivation, Lacrimation, Urination, Daefaction
- Also endothelial cells
 - Not innervated, can release NO

Nicotinic receptors

- Autonomic ganglia (Sympathetic & parasympathetic)
 - Fast responses Ligand-gated ion channel Action potential initiation

Skeletal neuromuscular junction

Nicotinic receptors

Fast responses – Ligand-gated ion channel
 Skeletal muscle contraction

Adrenoceptor localisation & action

Golan et al (Ed); Ch 10, p132

α -adrenoceptors

Blood vessels - constrict

Pupil (dilates) - constrict radial muscle

GIT - constrict sphincters

β -adrenoceptors

Heart - increase rate & force

Kidney - renin secretion

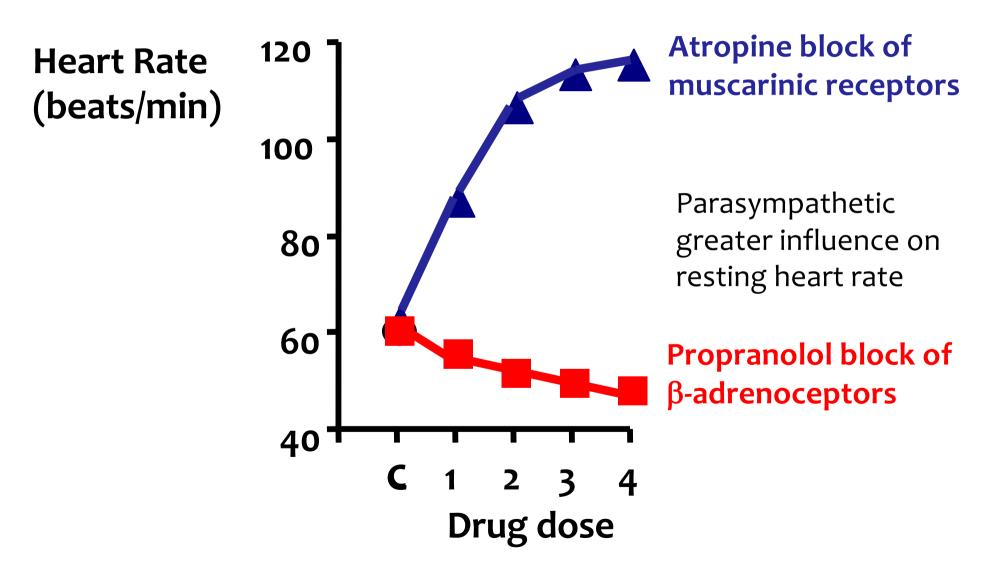
Skeletal BV's - dilate

Bronchi - dilate } (circulating Adr)

Autonomic Nervous System: Summary

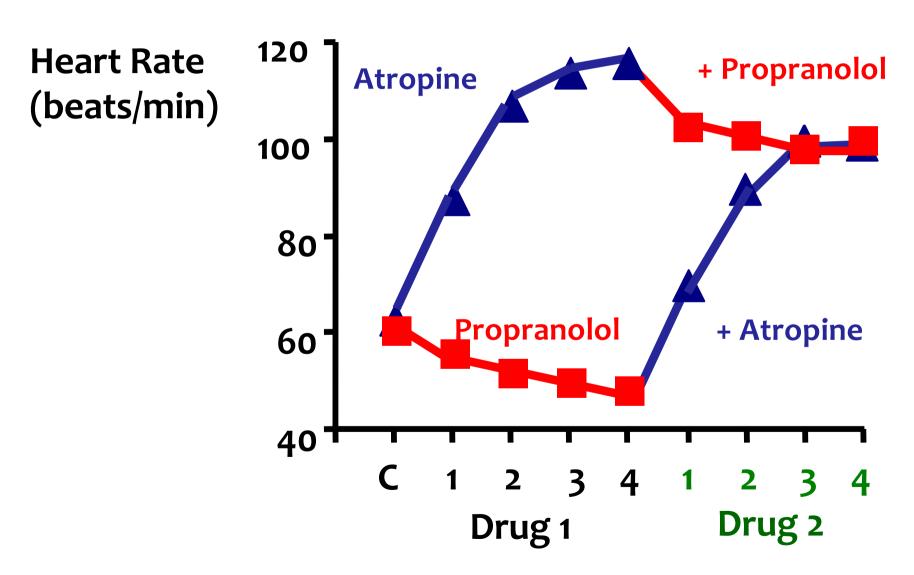
- Contrasting effects of sympathetic and parasympathetic nerves in tissues due to different transmitters
- Discrete responses for each division
 - Parasympathetic: acetylcholine, rest & repose responses
 - Sympathetic: noradrenaline, fight or flight responses
 - » (also involving circulating Adr release)
 - Receptors may be present in tissues that are not directly innervated
 - Muscarinic receptors on endothelium
 - $-\beta$ -adrenoceptors in airways
- In tissues receiving dual innervation
 - relative activity depends on needs of organ/tissue
- Nerves modulate activity
 - Pacemaker activity in heart, GIT motility

Autonomic control of heart rate at rest



Berne & Levy, Physiology 5th Ed. p323

Effect of combined autonomic blockade on resting heart rate

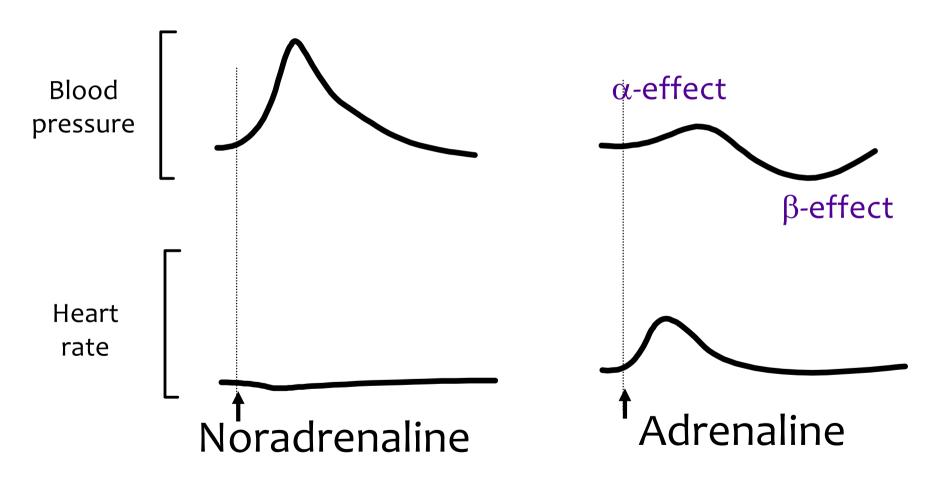


At the end of this lecture, you should be able to:

- •Describe the basic anatomy of the Peripheral NS (efferent), the Autonomic nervous System (ANS) and its Parasympathetic and Sympathetic divisions
- •Describe the major neurotransmitters and receptors involved in chemical transmission within the ANS and somatic NS
- List examples of agonists and antagonists that have selective activity at adrenoceptors

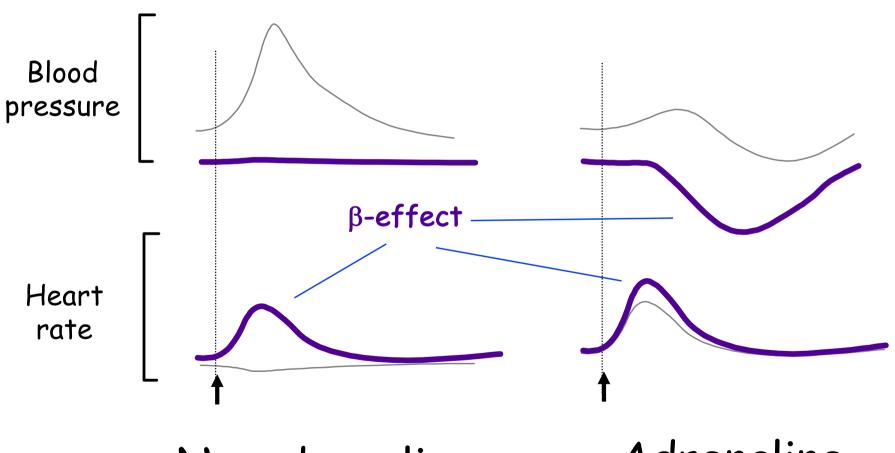
Cardiovascular adrenoceptor responses

Comparison of i.v. noradrenaline & adrenaline



Noradrenaline blood pressure increase can trigger reflex slowing of heart Direct β -effect can be offset by increased parasympathetic activity

Cardiovascular adrenoceptor responses Selective inhibition of α -effect by phentolamine



Noradrenaline Adrenaline Selective inhibition of β -effect by propranolol

α - and β - adrenoceptors

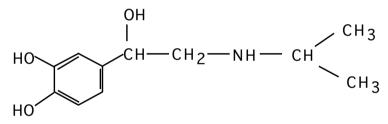
- Subtle differences between Noradrenaline and Adrenaline
 - Agonist receptor selectivity
 - NA \geq Adr at α -adrenoceptors
 - Adr \geq NA at β -adrenoceptors
- Selective inhibition of
 - ·BP increase by phentolamine
 - ·BP decrease and HR increase by propranolol
 - Antagonist receptor selectivity
 - Phentolamine α adrenoceptors
 - Propranolol β- adrenoceptors

Synthetic agonists with adrenoceptor selectivity

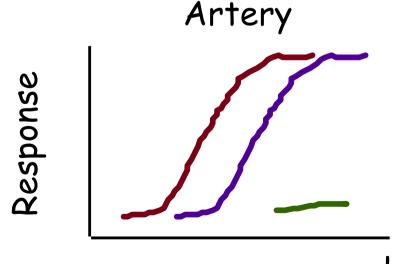
Noradrenaline

Phenylephrine

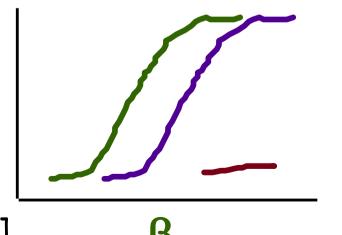
Isoprenaline



Heart



 α



log [agonist]

Therapeutic use of drugs with selectivity for β -adrenoceptors over α -adrenoceptors

Agonist: Isoprenaline

- Aerosol trialled in asthmatics
 - airway benefit offset by cardiac effects
 - » Cardiac palpitations
 - Salbutamol now used for bronchodilation
 - » Selectivity for airways, partial agonist

Antagonist: Propranolol

- Reduce cardiac output for hypertension
 - cardiac benefit offset by airway effect
- Still used clinically, but not if patient has asthma
 - » specific contraindications



(Laurence &Black, 1980)

"To get relief you must take some risk"