Circulation/Transport **General**

two major transport systems in body:

- A. The Circulatory System
- B. The Lymphatic Sysem

circulatory system works in conjunction with lymphatic system

→ they are directly connected to each other

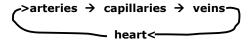
A. Circulatory (cardiovascular) System

circulatory system consists of "plumbing" and "pumps" & circulating fluid

pump = the heart

fluid = **blood**

blood flows in closed system of vessels over 60,000 miles of vessels (mainly capillaries)



arteries & arterioles

- take blood away from heart to capillaries

1

3

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

capillaries

B. Lymphatic System

fluid = lymph

venules & veins

-actual site of exchange

move lymph along

- bring blood from capillaries back to heart

an open system that returns excess materials

in the tissue spaces back to the blood

no dedicated pump; muscle contractions

direction; lymph does not circulate

lymphatic vessels move lymph in one

2

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

The Circulatory System (Cardiovascular System)

major connection between external and internal environment:

everything going in or out of body must go the circulatory system to get to where through its going

more than 60,000 miles of blood vessels with a pump that beats 100,000 times each day

General Functions of Circulatory System:

- A. Transport
- **B.** Homeostasis
- C. Protection

A. <u>Transport functions:</u>

- 1. Pick up food and oxygen from digestive and respiratory systems and deliver them to cells
- 2. pick up wastes and carbon dioxide from cells and deliver to kidneys and lungs
- 3. Transport hormones & other chemicals, enzymes etc throughout the body

B. Homeostasis functions:

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

- 4. maintain fluid and electrolyte balances in tissues and cells
- 5. maintain acid/base balances in tissues and cells
- 6. help regulate temperature homeostasis transfers excess heat from core to skin for removal

C. Protective Functions:

- 7. Clotting and Inflammation prevent excessive fluid loss and limit the spread of infection
- 8. Circulating cells and chemicals actively seek out and remove pathogens from the body
 - = "immune system"

The Heart - Anatomy

we are more aware of our heart than most other internal organs

Some ancient Chinese, Egyptian, Greek and Roman scholars correctly surmised that the heart is a pump for filling vessels with blood

Aristotle however thought the heart was the seat of emotion and a source of heat to aid digestion:

excited → heart beats faster "heartache" of grief

his thoughts predominated for over 2000 years before its true nature reemerged $\,$

the heart is one of first organ systems to appear in developing embryo

→ heart is beating by 4th week

study of heart = cardiology

no machine works as long or as hard as your heart

beats: >100,000 x's/day

> 30 Million times each year

> 3 Billion times in a lifetime to pump > 1 Million

barrels of blood

heart is about size and shape of closed fist

heart lies behind sternum in mediastinum,

broad superior border of heart = base

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

lower border of heart (=apex) lies on diaphragm

heart is enclosed in its own sac, = **pericardium** (=**pericardial sac**)(parietal pericardium) composed of tough fibrous outer layer and inner serous membrane

outer surface of heart is also covered with serous membrane (= visceral pericardium) (=epicardium) continuous with the pericardium

between the 2 membranes is **pericardial fluid**→lubrication

pericarditis = inflammation of pericardium,
 membranes become dry, each heartbeat becomes
 painful

wall of heart:

epicardium = visceral pericardium

thin & transparent serous tissue

myocardium = cardiac muscle cell most of heart

branching, interlacing contractile tissue acts as single unit (gap junctions)

endocardium = delicate layer of endothelial cells

continuous with inner lining of blood vessels [endocarditis]

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

6

Heart Chambers

interior of heart is subdivided into 4 chambers:

atria = two upper chambers
 with auricles
 smaller, thinner, weaker

ventricles = two lower chambers
larger, thicker, stronger
left ventricle much larger and thicker than
right ventricle

left ventricle is at apex of heart

Heart Vessels

There are 4 major vessels attached to heart:

2 arteries (take blood away from heart):

aorta

- from left ventricle

pulmonary trunk

- from right ventricle

2 veins (bring blood back to heart):

vena cava (superior & inferior)

- to right atrium

pulmonary veins (4 in humans)

- to left atrium

Heart Valves

There are also 4 one-way valves that direct flow of blood through the heart in one direction:

2 Atrioventricular (AV) valves

bicuspid (Mitral) valve

- separates left atrium and ventricle
- consists of two flaps of tissues

tricuspid valve

- separates right atrium and ventricle
- consists of three flaps of tissues

both held in place by chordae tendinae

attached to papillary muscles

→ prevent backflow (eversion) keeps valves pointed in direction of flow

2 Semilunar valves

at beginning of arteries leaving the ventricles

aortic SL valve

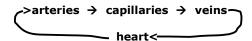
at beginning of aorta

pulmonary SL valve

at beginning of pulmonary trunk

Blood Vessels

blood flows in closed system of vessels over 60,000 miles of vessels (mainly capillaries)



arteries & arterioles

- take blood away from heart to capillaries

capillaries

-actual site of exchange

venules & veins

- bring blood from capillaries back to heart

Histology of Vessels

walls of arteries and veins consist of three layers:

- a. Tunica Externa
- b. Tunica Media
- c. Tunica Interna

a. Tunica Externa (= T. adventitia)

outer loose connective tissue

anchors the vessel and provides passage for small nerves, lymphatic vessels and smaller blood vessels

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

9

11

b. Tunica Media

middle, made mainly of smooth muscle with some elastic tissue and collagen fibers

strengthens vessel walls

→ prevent high pressure from rupturing them

allows vasodilation and vasoconstriction

usually the thickest layer, especially in arteries

c. Tunica Interna (=T. Intima)

inner endothelium

exposed to blood

when damaged or inflamed induce platelets or WBC's to adhere

→ may lead to plaque buildup and atherosclerosis

aneurysm = a weak point in arterial wall forms

Is a bulging sac that may rupture or put pressure on nearby brain tissue, vessels or other passageways.

usually due to degeneration of the tunica media, atherosclerosis or hypertension

Most common in abdominal aorta, renal arteries and circle of Willis

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

10

Types of Blood Vessels

1. Arteries & Arterioles

built to withstand the greatest pressure of the system

- →strong resilient walls,
- →thick layers of connective tissues
- →more muscular than veins

arteries and arterioles typically contain ~25% of all blood in circulation

15% in arteries; 10% in arterioles

pressure is variable

MAP ~ 93 varies from 100 - 40 mmHg

most organs receive blood from >1 arterial branch provides alternate pathways

2. Veins & Venules

generally have a greater diameter than arteries but thinner walls, flaccid

→ more compliant

three layer are all thinner than in arteries tunica adventitia is thickest of three

but not as elastic as arteries

little smooth muscle

 \sim 70% of all blood is in veins & venules \sim 60% in veins, \sim 10% in venules

low pressure:

12 – 8 mmHg venules 6 – 1 mmHg veins larger veins near 0

many of the medium veins, especially in limbs have = 1 way **valves**

3. Capillaries:

actual site of exchange of materials

→ the rest is just pumps and plumbing

consist of only a single layer of squamous epithelium= endothelial layer (=tunica intima)

arranged into capillary beds

= functional units of circulatory system

capillaries are extremely abundant in almost every tissue of the body

→ most of the 62,000 miles of blood vessels is capillaries

only 5% of blood at any one time is in capillaries

Circuits of Bloodflow

arteries, capillaries and veins are arranged into two circuits:

pulmonary: heart → lungs → heart

rt ventricle→ pulmonary arteries (trunk)→lungs→pulmonary veins→left atrium

systemic: heart → rest of body → heart left ventricle→aorta→body→vena cava→rt atrium

heart is a double pump

oxygen deficient blood in pulmonary artery and vena cava → usually blue on models

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

common iliac v. internal iliac v. external iliac v.

femoral v.

Anatomy of Circulatory System

Major Arteries and Veins

Pulmonary Circuit:

pulmonary a. Arteries

pulmonary v.

Systemic Circuit: Arteries

Veins:

13

15

ascending aorta rt & lft coronary a.

aortic arch brachiocephalic a.

on carotid a. internal carotid a. external carotid a. subclavian a.

axillary a. brachial a. Ift common carotid a.

lft subclavian a

descending aorta celiac trunk superior mesenteric a.

renal a. gonadal a. inferior mesenteric a.

common iliac a.

internal iliac a. external iliac a. femoral a.

superior vena cava

coronary v. brachiocephalic v. jugular v. subclavian v.

axillary v brachial v.

hepatic v.

hepatic portal v. esenteric v.

inferior mesenteric v. renal v.

gonadal v.

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

14

Special Circulation Patterns

1. Coronary Circulation (or Cardiac Circulation)

heart needs an abundant supply of oxygen and nutrients

→myocardium has its own supply of vessels

~5% of blood goes to heart muscle tissue ~10-x's its "fair share" based on weight alone

any interruption of blood flow can cause necrosis within minutes

= myocardial infarction

R & L Coronary Artery branch from aorta just beyond aortic SL valve

blood enters when Left Ventricle relaxes (most vessels receive blood when ventricles contract)

most blood returns to heart through veins that drain into coronary sinus

which empties into Right Atrium beneath entrance of Inferior Vena Cava

2. Circle of Willis

7 separate arteries

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

branchig from the internal carotids and vertebral arteries

arterial anastomosis interconnects them to form a circle of connecting arteries at base of brain

→ more than one route for blood to get to brain

3. Hepatic Portal System

veins from spleen, stomach, pancreas, gall bladder, and intestines

superior and inferior mesenteric merge to form hepatic portal vein

do not take blood directly to vena cava

instead take it to liver for "inspection"
-phagocytic cells remove toxins
-vitamins and minerals are stored

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

17

Heart Physiology

for the heart to work properly contraction and relaxation of chambers must be coordinated

Histology of Heart

cardiac muscle fibers

relatively short, thick branched cells

striated → myofibrils are highly ordered

usually 1 nucleus per cell

rather than tapering cells are bluntly attached to each other by gap junctions = **intercalated discs**

→ myocardium behaves as single unit

but atrial muscles separated from ventricular muscles by conducting tissue sheath

→ atria contract separately from ventricles

cardiac muscle cells cannot stop contracting to build up glycogen stores for anaerobic metabolism

→ need constant supply of oxygen & nutrients to

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

18

remain aerobic

→ greater dependence on oxygen than skeletal muscles

have exceptionally large mitochondria comprise 25% of cell volume (vs skeletal mm→2%)

cells are more adaptable in nutrient use; can use:

glucose fatty acids (preferred) lactic acid

Conducting System

cardiac muscle cells are not individually innervated as are skeletal muscle cells

→they are self stimulating

the rhythmic beating of the heart is coordinated and maintained by the heart conducting system

conducting system consists of:

SA Node

intrinsic rhythm 70-75 beats/min initiates stimulus that causes atria to contract (but not ventricles directly due to separation)

AV Node

picks up stimulus from SA Node if SA Node is not functioning it can act as a pacemaker =ectopic pacekmaker (usually slower intrinsic rhythm)

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

AV Bundle (Bundle of His)

connected to AV Node takes stimulus from AV Node to ventricles

Purkinje Fibers

takes impulse from AV Bundle out to cardiac mucscle fibers of ventricles causing ventricles to contract

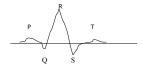
the heart conducting system generates a small electrical current that can be picked up by an electrocardiograph

=electrocardiogram (ECG; EKG)

ECG is a record of the electrical activity of the conducting system

body is a good conductor of electricity (lots of salts) potential changes at body's surface are picked up by 12 leads

[ECG is **NOT** a record of heart contractions]



P wave

passage of current through atria

from SA Node

conduction through atria is very rapid

atrial depolarization

QRS wave

passage of current through ventricles from AV Node – AV Bundle –

Purkinje Fibers

impulse slows as it passes to ventricles ventricular depolarization

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

T wave

repolarization of ventricles (atrial repolarization is masked by QRS)

by comparing voltage **amplitudes** and **time intervals**

between these waves from several leads can get idea of how rapidly the impulses are being conducted and how the heart is functioning

Abnormalities of ECG's = arrhythmias

1. bradycardia (<60 bpm)

decrease in body temperature some drugs (eg digitalis) overactive parasympathetic system endurance athletes

2. tachycardia (>100 bpm)

increased body temperature → fever emergencies, stress activation of sympathetic NS some drugs may promote fibrillation

3. flutter

short bursts of 200-300 bpm but coordinated

4. fibrillation

rapid, uncoordinated contractions of individual muscle cells atrial fibrillation is OK (since it only contributes 20% of blood to heart beat) ventricular fibrillation is lethal electrical shock used to defibrillate and recoordinate contractions

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

21

23

5. AV Node Block

normal P - Q interval = 0.12 - 0.20 seconds changes indicate damage to AV Node
→ difficulty in signal getting past AV Node

Cardiac Cycle

1 complete heartbeat (takes ~ 0.8 seconds)

consists of:

systole → contraction of each chamber

diastole → relaxation of each chamber

two atria contract simultaneously

as they relax, ventricles contract

relation of ECG to cardiac cycle

contraction and relaxation of ventricles produces characteristic **heart sounds**: lub-dub

abnormal sounds: "murmurs"

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

22

→ defective valves

congenital rheumatic (strep antibodies) septal defects

Cardiac Output

=The amount of blood that the heart pumps/min

CO = Heart Rate X Stroke volume = 75b/m X 70ml/b = 5250 ml/min (=5.25 l/min = ~1 gallon/min)

= ~ normal blood volume

in a lifetime the heart will pump ${\sim}53$ million gallons (200 Million L) of blood

during strenuous exercise heart may increase output 4 or 5 times this amount

A. Heart Rate:

innervated by **autonomic** branches to SA and AV nodes (antagonistic controls)

cardiac control center in medulla (cardiac center)

receives sensory info from:

Baroreceptors (stretch)

in aorta and carotid sinus increased stretch → slower

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

Chemoreceptors

monitor carbon dioxide and pH more CO₂ or lower pH → faster

B. Stroke Volume:

→normal SV = ~70 ml

(healthy heart pumps ~60% of blood in it)

also each side of heart must pump exactly the same amount of blood with each beat

→ otherwise excess blood would accumulate in lungs or in systemic vessels

eg. if Rt heart pumped 1 ml more per beat

→ within 90 minutes the entire blood volume
would accumulate in the lungs

most affected by:

mean arteriole pressure

systemic blood pressure = back pressure

condition of heart tissue

eg. heart contractility, fibrosis indicates amt of damage

Physiology of Blood Vessels

Blood circulates in arteries and capillaries by going down a **pressure gradient**

Blood Pressure

=the force of the blood flowing through blood vessels

measured as mmHg [100 mm Hg = 2 psi, tire $\sim 35 \text{psi}$]

changes in pressure are the driving force that moves blood through the circulatory system

blood pressure is created by

1. the force of the heart beat

previously discussed

the heart maintains a high pressure on the arterial end of the circuit

2. peripheral resistance

→ back pressure, resistance to flow

mainly depends on **diameter** of a vessel and its **compliance**

25

27

eg. **vasoconstriction** raises blood pressure

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

vasodilation lowers blood pressure

eg. **obesity** leads to many additional vessels that blood must pass through → raises blood pressure

[1lb of fat requires ~7 miles of blood vessels]

eg. any blockage of the normal diameter of a vessel will increase resistance

eg **atherosclerosis** inhibits flow \Rightarrow raises blood pressure

Measuring Blood Pressure

use sphygmomanometer

usually measure pressure in the brachial artery

procedure:

- a. increase pressure above systolic to completely cut off blood flow in artery
- b. gradually release pressure until 1st spurt (pulse) passes through cuff
 - = systolic pressure
- c. continue to release until there is no obstruction of flow sounds disappear

= diastolic pressure

normal BP = 120/80

range: 110-140 / 75-80 [mm Hg]

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

26

top number = systolic pressure; force of ventricular contraction

bottom number = diastolic pressure;
resistance of blood flow

may be more important

indicates strain to which vessels are continuously subjected

also reflects condition of peripheral vessels

Abnormal Blood Pressure

Hypotension = low BP: systolic <100:

usually not a cause for concern

often associated with long healthy life

but. in some may produce dizziness when standing up too quickly (esp in older patients)

may be due to severe bleeding and lead to circulatory shock

may hint at poor nutrition eg. <blood proteins

Hypertension = 140/90

if transient is normal: adaptation during fever, exercise, strong emotions

if persistent is a cause for concern

30% of those >50 yrs old suffer from hypertension

usually asymptomatic for first 10-20 yrs = silent killer

prolonged hypertension is a major cause of: heart failure, vascular disease, kidney failurel stroke, aneuryisms

high blood pressure affected by: heredity, gender (men at slightly higher risk of HBP), age (risk increases after age 35), race (African Americans at higher risk)

Flow of Blood in Veins

the blood pressure gradient is the main force that moves blood through arteries, arterioles and capillaries

movement of blood through veins is not pressure driven by the heart

veins have 1 - way valves

prevent backflow

most abundant in veins of limbs

quiet standing can cause blood to pool in veins and may cause fainting

varicose veins: "incompetent" valves

esp. superficial veins

may be due to; heredity, prolonged standing, obesity, pregnancy

hemorrhoids:

varicosities of anal veins

due to excessive pressure from birthing or bowel movements

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

venous pumps

muscular pump (=skeletal muscle pump)

during contraction veins running thru muscle are compressed and force blood in one direction (toward heart)

respiratory pump

inspiration: increases pressure in abdominopelvic cavity to push blood into thoracic cavity

expiration: increasing pressure in chest cavity forces thoracic blood toward heart

veins also act as blood reservoirs

 \Rightarrow with large lumens and thin walls they are compliant and can accommodate relatively large volumes of blood.

(60-70% of all blood is in veins at any time)

most organs are drained by >1 vein

- → occlusion of veins rarely blocks blood flow as it does in arteries
- → removal of veins during bypass surgery usually not traumatic

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

29

II. <u>Blood Flow & Differential Distribution of</u> Blood

the overall flow of blood to and within a particular organ or tissue is related to blood pressure and peripheral resistance

circulation also involves the **differential distribution** of blood to various body regions according to individual needs

→active body parts receive more blood than inactive parts

→blood volume must be shifted to parts as they become more active

these shifts are regulated by Vasomotor System

blood circulates because of pressure gradients

individual arterioles can increase or decrease their resistance to blood flow by constricting or dilating

mediated by autonomic nervous sytem

vasomotor control center in medulla

works in conjunction with cardiac centers

Capillaries & Capillary Beds

capillaries are the actual site of exchange of materials

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

20

 \rightarrow the rest is pumps and plumbing

each capillary <1mm long

thin walled - single cell layer thick

extremely abundant in almost every tissue of body

→most of 62,000 miles of vessels

usually no cell >0.1 mm away from a capillary

but only contains ~5% of blood in body

variable pressure 35 - 15 mm Hg

blood flows slowest in capillaries

due to greater **cross-sectional area** of all capillaries combined

ightarrow blood flows 1000x's faster in aorta than in capillaries

provides greatest opportunity for exchange to occur

Capillary Beds

capillary beds are the functional units of circulatory system

usually capillaries 10 –100 capillaries are organized into each capillary bed

arterioles and venules are joined directly by **metarterioles** (thoroughfare channels)

capillaries branch from metarterioles

cuff of smooth muscle surrounds origin of capillary branches

= precapillary sphincter

amount of blood entering a bed is regulated by:

- → vasomotor nerve fibers
- → local chemical conditions

31

Effects of Aging on CV System

most noticeable effect of aging on CV system is stiffening of arteries

heart has to work harder to overcome resistance

ventricles enlarge, esp left ventricle may get so thick that not enough space to pump blood effectively

valves may thicken and become calcified

impulse conduction along conducting system becomes more difficult

→increase in arrhythmias or heart block

muscle cells die

heart becomes weaker

→lower tolerance to physical activity

Atherosclerosis is main change seen in blood vessels with age

stiffening of arterial walls with increasing deposits of collagen fibers & declining resilience of elastic fibers

also decline in responsiveness of baroreceptors so less vasomotor response to changes in blood pressure

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

33

35

results: quick move from lying to standing, blood is drawn away from brain, can cause dizziness or fainting

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

34

Disorders of the Circulatory System

Heart Disease

can lead to heart attack and ultimately heart failure

leading cause of death in US for both men and women \rightarrow 500,000 deaths/yr 7.2 M/yr worldwide (°07) \rightarrow 30% of deaths/yr

most common form is **coronary atherosclerosis**often leading to myocardial infarction (heart attack)

Heart Attack

heart attack risk is ~'50% genetic & 50% cheeseburger'

begins with the buildup of plaque:

- a. cholesterol in blood infiltrates the arterial wall
- b. immune system dispatches macrophages to consume the
- c. macrophages become foam cells full of cholesterol
- d. foam cells accumulate and become a major component of plaque
- e. to keep the arterial wall slick, smooth muscle cells form a $\ensuremath{\mathsf{cap}}$
- f. foam cells in plaque secrete chemicals that weaken the $\ensuremath{\mathsf{cap}}$
- g. f the cap cracks, plaque seeps into the blood stream and a clot forms that blocks bloodflow

Abnormal Blood Pressure

Hypotension

low BP → systolic <100
usually not a cause for concern
→ often associated with long healthy life

but.

in some may produce dizziness when standing up too quickly (esp in older patients) may be due to severe bleeding and lead to circulatory shock may hint at poor nutrition eg.

eg.
blood proteins

Hypertension

if transient is normal:

adaptation during fever, exercise, strong emotions

if persistent is a cause for concern (silent killer) 30% of those >50 yrs old suffer from hypertension usually asymptomatic for first 10-20 yrs= silent killer

high blood pressure affected by:

gender: men slightly higher risk of HBP age: risk increases after age 35 heredity/race: African Americans at higher risk

diet: any factors leading to obesity

prolonged hypertension is a major cause of: heart failure vascular disease kidney failure stroke aneuryisms

<u>Stroke</u>

sudden death of brain tissue occurring when

cerebral atherosclerosis, thrombosis or hemorrhage of a cerebral aneurysm cuts off blood flow to part of the brain.

effects range from unnoticeable to fatal depending on extent of tissue damage and function of affected tissue

Varicose Veins

can occur anywhere on body but most common on legs veins in legs are largest in body and must counteract gravity

to get blood back to the heart

veins become enlarged and valves fail to prevent backflow of blood

often associated with tired, achy, or feeling of heavy limbs most common in superficial saphenous veins

→ they are poorly supported by surrounding tissues

many factors contribute to likelihood of varicose veins:

heredity

age esp occur betw 18 and 35 yrs, peaks betw 50 and 60

gender women are 4 to 1 times more likely to get them **pregnancy** sometimes form during pregnancy (8-20% chance) then disappear afterwards

lifestyle: prolonged sitting or standing daily

Transposition of the Great Vessels

the child will develop normally until they begin to walk the right ventricle wll be unable to pump enough blood through systemic circuit

Human Anatomy & Physiology: Circulatory System, Ziser Lecture Notes, 2013.11

37