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CELLULAR ACTIVITIES, STEPS, HORMONAL REGULATION, CLINICAL SIGNIFICANCE, DIAGNOSIS AND TREATMENT OF BONE REMODELLING

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ABSTRACT:-

Osteoclasts, osteoblasts and osteocytes are seen in the bone. Osteoclasts play a role in bone resorption. Osteoclasts release enzymes and acids that attack the mineralized matrix of bone, releasing calcium and other minerals into the blood stream. Osteoblasts play a role in the synthesis and deposition of collagen and other protein which behave as a scaffold for the mineralization of new bone tissue. Osteocytes play a role in finding mechanical stress amd controlling bone remodelling by signalling to osteoclasts as well as osteoblasts. Bone remodelling is controlled by hormones such as calcitonin (CT) and Vitamin D. Cytikines like RANKL and OPG are also involved in the regulation of osteoclast. Growth factors show impact on bone healing as well as remodelling. Osteoporosis and Pagets disease are treated as bone disorders. Clinical significance of bone remodelling include maintenance of bone density, fracture healing and calcium homeostasis.

KEY WORDS: Mineralized matrix, enzymes, acids, calcium, collagen, mechanical stresses, bone remodelling units (BRU), osteoid, fracture remodelling, Para thyroid hormone (PTH), Calcitonin (CT), vitamin D, Cytokines, growth factors, osteoporosis, pagets disease, bone density, calcium homeostasis, osteoporosis, biphosphonates, orthopedic surgery, cancer meta stasis and aging.

Bone remodeling is a dynamic and continuous process that occurs throughout your life. It involves the removal of old bone tissue (resorption) and the formation of new bone tissue (ossification) to maintain the integrity and strength of your skeletal system. This process is essential for various functions, including repair of fractures, calcium homeostasis, and adaptation to mechanical stress.

ISSN- 2394-5125 VOL 10, ISSUE 07, 2023

CELLULAR ACTIVITIES IN BONE REMODELLING:-

Osteoclasts: These are specialized cells responsible for bone resorption. They secrete enzymes and acids that break down the mineralized matrix of bone, releasing calcium and other minerals into the bloodstream.

Osteoblasts: Osteoblasts are bone-forming cells. They synthesize and deposit collagen and other proteins, which serve as a scaffold for the mineralization of new bone tissue.

Osteocytes: These are mature osteoblasts that have become embedded within the bone matrix. Osteocytes play a crucial role in detecting mechanical stresses and regulating bone remodeling by signaling to osteoclasts and osteoblasts.

STEPS IN BONE REMODELLING:-

Activation: The process begins with the activation of bone remodeling units (BRUs) at specific sites within bone tissue. Various factors, including hormones, mechanical stress, and microdamage, trigger this activation.

Resorption: Osteoclasts are recruited to the activated site. They create a resorption pit by breaking down old bone tissue. This releases calcium and phosphate into the bloodstream.

Reversal: After resorption, the bone undergoes a reversal phase during which a transition from resorption to formation occurs. Osteoclasts are replaced by osteoblasts.

Formation: Osteoblasts move to the resorbed area and begin laying down new bone tissue. Initially, they produce an unmineralized matrix called osteoid.

Mineralization: Over time, the osteoid becomes mineralized as calcium and phosphate are deposited, forming mature bone tissue.

Quiescence: Finally, some osteoblasts become osteocytes, which remain embedded in the bone matrix and help regulate future remodeling.

REGULATION OF BONE REMODELLING:-

Hormonal Regulation

Parathyroid Hormone (PTH): Effects on calcium and phosphate balance.

Calcitonin: Role in calcium regulation.

Vitamin D: Importance in bone health.

Mechanical Forces and Bone Remodeling

Wolff's Law: Adaptation of bone to mechanical stress.

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Importance of weight-bearing exercise.

Cytokines and Growth Factors

Role of cytokines like RANKL and OPG in osteoclast regulation.

Growth factors' influence on bone healing and remodeling.

Aging and Bone Remodeling

Changes in bone remodeling with age.

Risk factors for osteoporosis.

Diseases Affecting Bone Remodeling

Osteoporosis: Pathogenesis and treatment.

Paget's Disease: Impact on bone turnover.

Pharmacological Interventions

Medications like bisphosphonates and RANKL inhibitors.

Their mechanisms and clinical use.

Nutritional Factors

Role of calcium, vitamin D, and other nutrients in bone health.

Dietary recommendations for maintaining strong bones.

Age and Health: Bone remodeling rates change with age, and certain medical conditions can affect the process, leading to bone diseases like osteoporosis.

CLINICAL SIGNIFICANCE:-

Bone remodeling is a crucial physiological process in the human body with several clinical significances:

Maintenance of Bone Density: It helps maintain bone density by removing old or damaged bone tissue and replacing it with new, healthy bone, preventing osteoporosis and fractures.

Fracture Healing: After a bone fracture, remodeling is essential for the healing process. It helps restore the bone's strength and shape.

Calcium Homeostasis: Bone remodeling regulates calcium levels in the blood, which is vital for various physiological functions. Imbalances can lead to conditions like hypercalcemia or hypocalcemia.

DISEASE DIAGNOSIS:-

ISSN- 2394-5125 VOL 10, ISSUE 07, 2023

Abnormal bone remodeling can indicate underlying health issues. For instance, increased bone resorption may be a sign of osteoporosis, while decreased remodeling can be seen in conditions like osteopetrosis.

DRUG DEVELOPMENT:-

Understanding bone remodeling is crucial for developing medications to treat bone disorders. Drugs like bisphosphonates target this process to manage conditions like osteoporosis.

Orthopedic Surgery: Surgeons consider bone remodeling when planning procedures like joint replacements. It affects the long-term success and stability of implants.

Cancer Metastasis: Bone remodeling can be affected by certain cancers, such as metastatic breast cancer, which can lead to bone pain and fractures.

Aging: Age-related changes in bone remodeling can contribute to the risk of fractures and osteoporosis in older individuals.

CONCLUSION:-

Bone remodeling plays a fundamental role in maintaining bone health and has far-reaching clinical implications in various medical fields, from orthopedics to endocrinology. Understanding this process is essential for diagnosing, treating, and preventing bone-related disorders.

REFERENCES

1. Frost HM. Wolff's Law and bone's structural adaptations to mechanical usage: an overview for clinicians. Angle Orthod. 1994;64(3):175-88. [PubMed]

2. Siddiqui JA, Partridge NC. Physiological Bone Remodeling: Systemic Regulation and Growth Factor Involvement. Physiology (Bethesda). 2016 May;31(3):233-45. [PMC free article] [PubMed]

3 .Delaisse JM. The reversal phase of the bone-remodeling cycle: cellular prerequisites for coupling resorption and formation. Bonekey Rep. 2014;3:561. [PMC free article] [PubMed]

3. Everts V, Delaissé JM, Korper W, Jansen DC, Tigchelaar-Gutter W, Saftig P, Beertsen W. The bone lining cell: its role in cleaning Howship's lacunae and initiating bone formation. J Bone Miner Res. 2002 Jan;17(1):77-90. [PubMed]

4. Schaffler MB, Cheung WY, Majeska R, Kennedy O. Osteocytes: master orchestrators of bone. Calcif Tissue Int. 2014 Jan;94(1):5-24. [PMC free article] [PubMed]

6 .Hadjidakis DJ, Androulakis II. Bone remodeling. Ann N Y Acad Sci. 2006 Dec;1092:385-96. [PubMed]

7.Wein MN, Kronenberg HM. Regulation of Bone Remodeling by Parathyroid Hormone. Cold Spring Harb Perspect Med. 2018 Aug 01;8(8) [PMC free article] [PubMed]

ISSN- 2394-5125 VOL 10, ISSUE 07, 2023

8. Khosla S, Oursler MJ, Monroe DG. Estrogen and the skeleton. Trends Endocrinol Metab. 2012 Nov;23(11):576-81. [PMC free article] [PubMed]

9. Olney RC. Regulation of bone mass by growth hormone. Med Pediatr Oncol. 2003 Sep;41(3):228-34. [PubMed]

10. Canalis E, Delany AM. Mechanisms of glucocorticoid action in bone. Ann N Y Acad Sci. 2002 Jun;966:73-81. [PubMed]

11. Bassett JH, Williams GR. Role of Thyroid Hormones in Skeletal Development and Bone Maintenance. Endocr Rev. 2016 Apr;37(2):135-87. [PMC free article] [PubMed]

12. Seibel MJ. Biochemical markers of bone turnover: part I: biochemistry and variability. Clin Biochem Rev. 2005 Nov;26(4):97-122. [PMC free article] [PubMed]

13. Garnero P. The Utility of Biomarkers in Osteoporosis Management. Mol Diagn Ther. 2017 Aug;21(4):401-418. [PubMed]

14.Adams C, Banks KP. StatPearls [Internet]. StatPearls Publishing; Treasure Island (FL): Aug 8, 2022. Bone Scan. [PubMed]

15. Kucukalic-Selimovic E, Begic A. Value of bone scintigraphy for detection and ageing of vertebral fractures in patients with severe osteoporosis and correlation between bone scintigraphy and mineral bone density. Med Arh. 2004;58(6):343-4. [PubMed]

16. de Vernejoul MC. Bone remodelling in osteoporosis. Clin Rheumatol. 1989 Jun;8 Suppl 2:13-5. [PubMed]

17. Christiansen P. The skeleton in primary hyperparathyroidism: a review focusing on bone remodeling, structure, mass, and fracture. APMIS Suppl. 2001;(102):1-52. [PubMed]

18. Sabharwal R, Gupta S, Sepolia S, Panigrahi R, Mohanty S, Subudhi SK, Kumar M. An Insight in to Paget's Disease of Bone. Niger J Surg. 2014 Jan;20(1):9-15. [PMC free article] [PubMed]

19. Stark Z, Savarirayan R. Osteopetrosis. Orphanet J Rare Dis. 2009 Feb 20;4:5. [PMC free article] [PubMed]

20. Ganesan K, Goyal A, Roane D. StatPearls [Internet]. StatPearls Publishing; Treasure Island (FL): Sep 5, 2022. Bisphosphonate. [PubMed]

21 .McLaughlin MB, Jialal I. StatPearls [Internet]. StatPearls Publishing; Treasure Island (FL): Jul 18, 2022. Calcitonin. [PubMed]

22.Quintanilla Rodriguez BS, Correa R. StatPearls [Internet]. StatPearls Publishing; Treasure Island (FL): Feb 13, 2023. Raloxifene. [PubMed]