

Nanotechnology And Its Applications In Orthopaedics

Dr. Nilargha Chowdhury

Post-Graduate Trainee, Department Of Orthopaedics, Pacific Institute Of Medical Sciences, Udaipur, India

Dr. Asif Husain Ansari

Junior Resident, Department Of Orthopaedics, Government Medical College, Bhilwara, India

Dr. Nisith Ranjan Chowdhury

Senior Consultant, Department Of Sports Medicine, Sebayam Nursing Home, West Bangal, India

Dr. Aabid Husain Ansari

Assistant Professor, Department Of Orthopaedics, Pacific Institute Of Medical Sciences, Udaipur, India

Abstract:

Nanotechnology has emerged as a groundbreaking field with transformative potential in various sectors, including orthopaedics. The integration of nanoscale materials and technologies into orthopaedic practices has opened up new avenues for advancing treatment modalities, implant design, and diagnostic capabilities. In this article, we delve into the role of nanotechnology in orthopaedics, exploring its applications in implant technology, bone regeneration, drug delivery systems, and imaging diagnostics. By harnessing the unique properties of nanomaterials, researchers and healthcare professionals are paving the way for innovative solutions to enhance patient outcomes and quality of care in orthopaedic medicine.

Keywords: Nanotechnology, Orthopedic implant, Nanoparticles, Nanomaterials, Nanocoating

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I. Introduction:

The term "nanotechnology" was first used by Dr. Norio Taniguchi, a professor at the Science University of Tokyo. Richard Feynman was the first to conceptualize the potential of nanotechnology in 1959 [1]. It generally refers to the branch of engineering that deals with objects having one dimension ranging from 1 to 100 nm [2]. A nanometer is defined as a unit of measure corresponding to 10^{-9} meter. Nanotechnology is the art of manipulating materials at the nanoscale, a size so small it makes a grain of sand look massive. It's all about manipulating things at the tiniest level to create big impacts. In the realm of orthopaedics, nanotechnology has emerged as a promising avenue for enhancing the diagnosis, treatment, and management of musculoskeletal disorders. In orthopaedics, this techy magic helps in creating materials and structures that interact seamlessly with the body's bones and tissues. From enhancing bone regeneration to enabling targeted drug delivery and improving the design of orthopaedic implants, nanotechnology offers innovative solutions that hold great promise for improve patient outcomes, enhance treatment efficacy, reduce recovery times, and improve overall quality of care for orthopaedic patients worldwide [3,4]. This article explores the pivotal role of nanotechnology in orthopaedics, delving into its diverse applications, potential benefits, and the future directions that hold promise for transforming orthopaedic care.

II. Nanotechnology And Its Applications In Orthopaedics

Nanotechnology, the science of manipulating matter at the nanoscale, has found exciting applications in orthopaedics. By utilizing nanomaterials and techniques, researchers aim to enhance bone repair, tissue regeneration, drug delivery systems, and implant development [Figure 1].

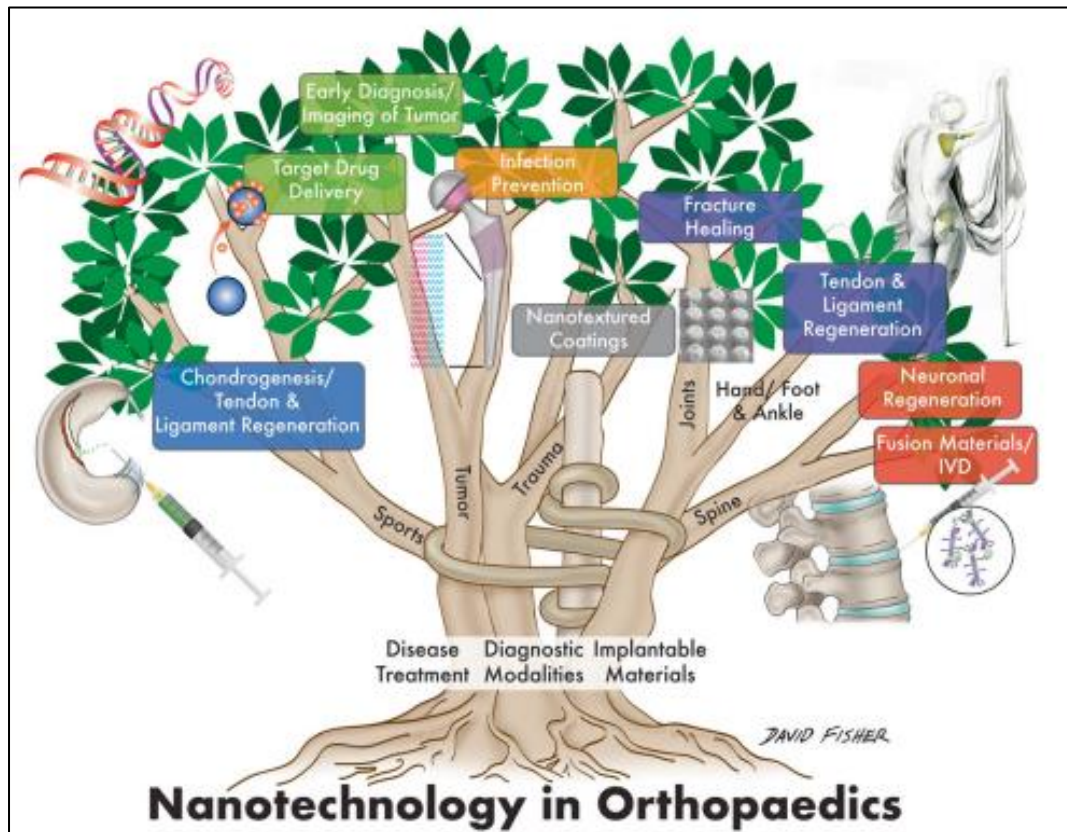


Figure 1. Diagram showing applications of Nanotechnology in Orthopaedics (Smith et al. 2018)².

Nanomaterials for Bone Regeneration and Repair:

Nanotechnology offers the ability to create nanomaterials and nanoparticles that play a crucial role in bone tissue engineering by providing a scaffold for cell growth and promoting bone regeneration. Their small size allows for better interaction with cells, leading to improved healing and tissue formation. Nanofibers and nanocomposites offer unique properties that make them ideal for bone regeneration applications. These materials mimic the natural structure of bone, providing mechanical support and promoting new bone formation for enhanced healing [5-7].

Nanofibrous Scaffolds for Tissue Engineering:

Nanofibrous scaffolds provide support and guidance for new tissue growth, helping in bone & tissue repair and regeneration. Nanofibers scaffolds encouraging bone cells to grow and build new tissue. Nanoparticles releases growth factors at just the right time and place, guiding bone regeneration like expert conductors leading a symphony. Poly (γ -glutamic acid) nanocomplexes injection has been shown in multiple studies to enhance the intervertebral disc regeneration [8-10]. Nanoscale synthetic nerve conduits like carbon nanotubes, have more tunable mechanical properties than autografts and have ability to promote axonal growth and may enhance nerve regeneration through augmented surface topographical interactions [11,12]. Nanofibrous scaffold composed of polycaprolactone, gelatin, polyethersulfone, hydrogels, and peptide based materials etc. can be used for Chondrogenesis. They enhanced articular cartilage repair by facilitating mesenchymal stem cells differentiation towards the proper chondrogenic lineage [13-15]. Nanocomposite scaffolds also helps in tendon and ligaments regeneration. These scaffolds facilitate improved healing and mechanical stability, and prevents tendon adhesion formation [16-17].

Nanoparticles for Targeted Drug Delivery in Orthopaedics:

Nanoparticles are used in targeted drug delivery systems for orthopaedic treatments of malignant bone tumors, prosthetic joint infections, and osteomyelitis. They allow precise delivery of therapeutic agents to the site of injury or inflammation. This targeted approach minimizes side effects and improves treatment outcomes. In treatment of malignant bone tumors like Osteosarcoma and Ewing sarcoma, chemotherapy drug loaded nanoparticles bind and enter the cancer cell and directly damaged the cancerous cells with reduced collateral toxicity to non-cancerous cells [18]. The addition of nanotechnology-based antibiotic carriers such as lipid nanoparticles [19], silica [20], and clay nanotubes [21] to bone cement may enhance drug delivery, allow for

timed release, increased osseointegration and osteoblast activity [22,23]. Nanophase silver has anti-microbial property and is used clinically in wound care. It decrease bacterial colonization and prevent infection [24,25].

While nanoparticle drug delivery offers many benefits, such as improved efficacy and reduced systemic toxicity, there are challenges to overcome, including ensuring proper distribution and clearance of nanoparticles. Research continues to address these issues for optimal treatment results.

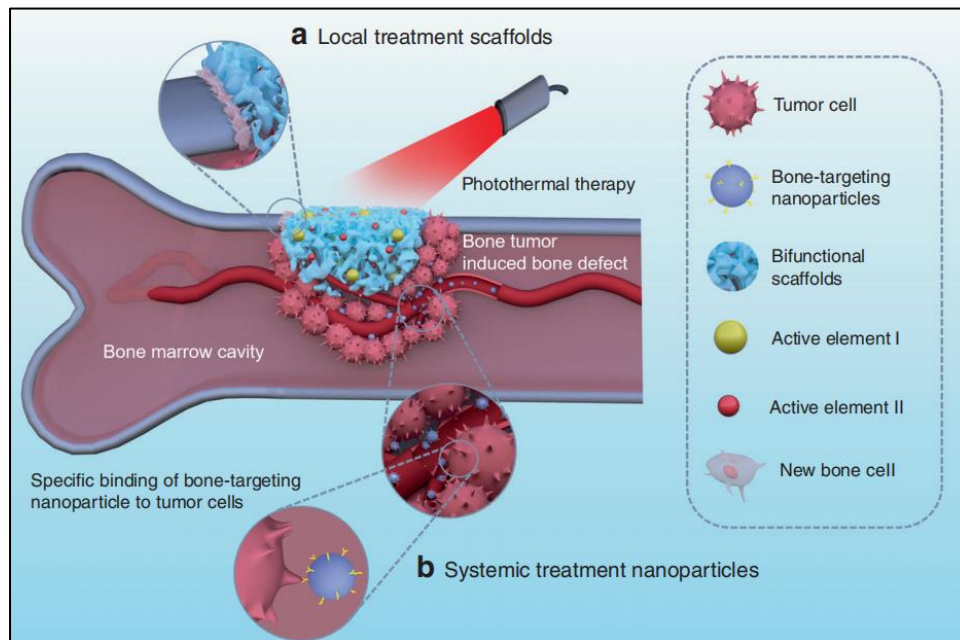


Figure 2. Bifunctional biomaterials include (A) local treatment scaffolds (such as 3D-printed scaffolds, nano/microparticle-containing scaffolds, and hydrogels) and (B) systemic treatment nanoparticles (like bone-targeting nanoparticles) for tumor photothermal therapy and bone regeneration (J Liao et al. 2021)²⁶.

Nanoengineering of Orthopaedic Implants:

Nanocoatings are applied to orthopaedic implants to improve their integration with surrounding bone tissue. These coatings enhance osseointegration, reduce implant rejection, and promote long-term stability for better patient outcomes [Figure 3]. Customized nanoscale surface modifications on orthopaedic implants can tailor their properties to specific patient needs. By fine-tuning surface characteristics at the nanoscale, implants can exhibit enhanced antibacterial properties, boost strength reduced wear and tear, and improved biomechanical performance and longevity. It's like giving implants a high-tech boost to ensure they stand the test of time in the bone battlefield.

Nanocoatings of titanium oxide and zirconia on titanium spinal implants promotes increased bone formation and decreased resorption compared to conventional smooth implants [27]. Silicon nitride nanoparticles coated cervical cages have demonstrated multiple biomechanical advantages over standard PEEK (poly-ether-ether-ketone) for interbody fusion [28]. Nano-selenium implants in bone cancer inhibit the growth of malignant osteoblasts while promoting healthy bone function at the implant-tissue interface, increased bone adhesion, calcium deposition, bone proliferation, and alkaline phosphatase activity [29]. Nanotextured implants improve the biocompatibility and mechanical properties of titanium implants and stimulate implant osseointegration and surrounding osteogenesis to a greater degree than conventional implants in replacement surgery [30]. The addition of carbon nanotubes to ultra-high molecular weight polyethylene (UHMWPE) liner, improve its biocompatibility properties, reduces wear and tear and reduces the risk of fracture of liner [31].

There are some challenges associated with using nanotechnology in orthopaedic implants include ensuring the biocompatibility of nanomaterials, addressing potential toxicity concerns, and overcoming regulatory hurdles related to the use of novel nano-engineered materials in medical devices. It's like navigating a nano-sized obstacle course for implant perfection.

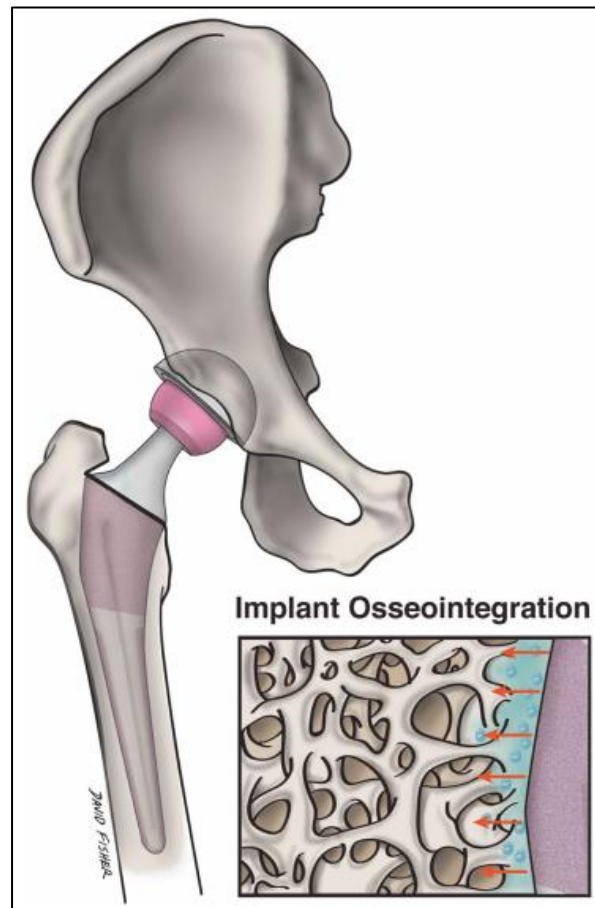


Figure 3. Diagram showing magnified nanoengineered implant surface and its topographical connection with the surrounding bone (Smith et al. 2018)².

Nanoscale Diagnostics and Imaging Techniques in Orthopaedics:

Nanotechnology enables the development of highly sensitive imaging techniques and nanosensors that can detect orthopaedic conditions at an early stage, leading to improved diagnostic accuracy and treatment outcomes. Nanoparticles are like the tiny detectives of the orthopaedic world, helping us see what's going on inside the body with greater clarity. These minuscule particles can be designed to seek out specific areas of concern, allowing for more precise imaging of orthopaedic conditions. They can detect changes at the molecular level, alerting surgeons to potential orthopaedic issues long before they become major problems [32-34]. For example in osteoarthritis or osteoporosis, by catching these issues sooner rather than later, patients can receive timely interventions to prevent further deterioration and improve their quality of life. By harnessing nanoscale imaging, orthopaedic researchers can explore the intricacies of bone structure, cartilage composition, and even implant integration at a whole new level. This technology isn't just about pretty pictures – it's about uncovering the mysteries of orthopaedic conditions and developing more precise diagnostic tools.

III. Future Directions And Challenges In Nanotechnology For Orthopaedics:

The future of nanotechnology in orthopaedics is brighter than a titanium hip implant. As researchers delve deeper into the nano realm, new possibilities emerge for improving orthopaedic treatments and outcomes. Nanotechnology has the potential to revolutionize orthopaedic surgery by enabling precision treatments, personalized implants, targeted drug delivery systems, and enhanced post-operative recovery, ultimately improving patient care and surgical outcomes. From nanoscale drug delivery systems to bioactive implants, the potential applications of nanotechnology in orthopaedics are expanding rapidly. But, as with any cutting-edge technology, there are challenges and ethical considerations to navigate. We must address challenges such as regulatory hurdles, safety concerns, and the ethical implications of manipulating materials at the nano level. Balancing innovation with responsibility is key to ensuring that nanotechnology benefits patients without unintended consequences.

IV. Conclusion:

In conclusion, the integration of nanotechnology into orthopaedic practices holds immense promise for revolutionizing the way we approach bone regeneration, implant development, and diagnostic imaging. As research advances and technologies mature, the potential for personalized and more effective orthopaedic treatments becomes increasingly tangible. By leveraging the principles of nanotechnology, orthopaedic practitioners can optimize patient care, improve surgical outcomes, and pave the way for a new era of precision medicine in orthopaedics.

Conflicts of interest:

The authors have none to declare.

References:

- [1] Feynman R. There's Plenty Of Room At The Bottom. *Infeynman And Computation* 2018 Mar 8 (Pp. 63-76). Crc Press.
- [2] Smith Wr, Hudson Pw, Ponce Ba, Rajaram Manoharan Sr. Nanotechnology In Orthopedics: A Clinically Oriented Review. *Bmc Musculoskeletal Disorders*. 2018 Dec;19:1-0.
- [3] Pleshko N, Grande Da, Myers Kr. Nanotechnology In Orthopaedics. *Jaaos-Journal Of The American Academy Of Orthopaedic Surgeons*. 2012 Jan 1;20(1):60-2.
- [4] Wong Kk, Liu Xl. Nanomedicine: A Primer For Surgeons. *Pediatric Surgery International*. 2012 Oct;28:943-51.
- [5] Balasundaram G, Webster Tj. An Overview Of Nano-Polymers For Orthopedic Applications. *Macromolecular Bioscience*. 2007 May 10;7(5):635-42.
- [6] Webster Tj, Ejiiofor Ju. Increased Osteoblast Adhesion On Nanophase Metals: Ti, Ti6al4v, And Cocrho. *Biomaterials*. 2004 Aug 1;25(19):4731-9.
- [7] Yi H, Ur Rehman F, Zhao C, Liu B, He N. Recent Advances In Nano Scaffolds For Bone Repair. *Bone Research*. 2016 Dec 13;4(1):1-1.
- [8] Antunes Jc, Pereira Cl, Teixeira Gq, Silva Rv, Caldeira J, Grad S, Gonçalves Rm, Barbosa Ma. Poly (Γ-Glutamic Acid) And Poly (Γ-Glutamic Acid)-Based Nanocomplexes Enhance Type Ii Collagen Production In Intervertebral Disc. *Journal Of Materials Science: Materials In Medicine*. 2017 Jan;28:1-7.
- [9] Teixeira Gq, Pereira Cl, Castro F, Ferreira Jr, Gomez-Lazaro M, Aguiar P, Barbosa Ma, Neidlinger-Wilke C, Goncalves Rm. Anti-Inflammatory Chitosan/Poly-Γ-Glutamic Acid Nanoparticles Control Inflammation While Remodeling Extracellular Matrix In Degenerated Intervertebral Disc. *Acta Biomaterialia*. 2016 Sep 15;42:168-79.
- [10] Cui X, Liu M, Wang J, Zhou Y, Xiang Q. Electrospun Scaffold Containing Tgf-B1 Promotes Human Mesenchymal Stem Cell Differentiation Towards A Nucleus Pulposus-Like Phenotype Under Hypoxia. *Iet Nanobiotechnology*. 2015 Apr;9(2):76-84.
- [11] Mattei Ta, Rehman Aa. "Extremely Minimally Invasive": Recent Advances In Nanotechnology Research And Future Applications In Neurosurgery. *Neurosurgical Review*. 2015 Jan;38:27-37.
- [12] Dalamagkas K, Tsintou M, Seifalian A. Advances In Peripheral Nervous System Regenerative Therapeutic Strategies: A Biomaterials Approach. *Materials Science And Engineering: C*. 2016 Aug 1;65:425-32.
- [13] Mahboudi H, Kazemi B, Soleimani M, Hanaee-Ahvaz H, Ghanbarian H, Bandehpour M, Enderami Se, Kehtari M, Barati G. Enhanced Chondrogenesis Of Human Bone Marrow Mesenchymal Stem Cell (Bmsc) On Nanofiber-Based Polyethersulfone (Pes) Scaffold. *Gene*. 2018 Feb 15;643:98-106.
- [14] Hastar N, Arslan E, Guler Mo, Tekinay Ab. Peptide-Based Materials For Cartilage Tissue Regeneration. *Peptides And Peptide-Based Biomaterials And Their Biomedical Applications*. 2017:155-66.
- [15] Liu J, Nie H, Xu Z, Niu X, Guo S, Yin J, Guo F, Li G, Wang Y, Zhang C. The Effect Of 3d Nanofibrous Scaffolds On The Chondrogenesis Of Induced Pluripotent Stem Cells And Their Application In Restoration Of Cartilage Defects. *Plos One*. 2014 Nov 12;9(11):E111566.
- [16] Huegel J, Kim Dh, Cirone Jm, Pardes Am, Morris Tr, Nuss Ca, Mauck Rl, Soslowsky Lj, Kuntz Af. Autologous Tendon-Derived Cell-Seeded Nanofibrous Scaffolds Improve Rotator Cuff Repair In An Age-Dependent Fashion. *Journal Of Orthopaedic Research*. 2017 Jun;35(6):1250-7.
- [17] Verdiyeva G, Koshy K, Glibbery N, Mann H, Seifalian Am. Tendon Reconstruction With Tissue Engineering Approach—A Review. *Journal Of Biomedical Nanotechnology*. 2015 Sep 1;11(9):1495-523.
- [18] Savvidou Od, Bolia Ik, Chloros Gd, Goumenos Sd, Sakellariou Vi, Galanis Ec, Papagelopoulos Pj. Applied Nanotechnology And Nanoscience In Orthopedic Oncology. *Orthopedics*. 2016 Sep 1;39(5):280-6.
- [19] Ayre Wn, Birchall Jc, Evans Sl, Denyer Sp. A Novel Liposomal Drug Delivery System For Pmma Bone Cements. *Journal Of Biomedical Materials Research Part B: Applied Biomaterials*. 2016 Nov;104(8):1510-24.
- [20] Shen Sc, Ng Wk, Dong Yc, Ng J, Tan Rb. Nanostructured Material Formulated Acrylic Bone Cements With Enhanced Drug Release. *Materials Science And Engineering: C*. 2016 Jan 1;58:233-41.
- [21] Wei W, Abdullayev E, Hollister A, Mills D, Lvov Ym. Clay Nanotube/Poly (Methyl Methacrylate) Bone Cement Composites With Sustained Antibiotic Release. *Macromolecular Materials And Engineering*. 2012 Jul;297(7):645-53.
- [22] No Yj, Roohani-Esfahani Si, Zreiqat H. Nanomaterials: The Next Step In Injectable Bone Cements. *Nanomedicine*. 2014 Aug;9(11):1745-64.
- [23] Ricker A, Liu-Snyder P, Webster Tj. The Influence Of Nano Mgo And Baso4 Particle Size Additives On Properties Of Pmma Bone Cement. *International Journal Of Nanomedicine*. 2008 Dec 1;3(1):125-32.
- [24] Jia Z, Xiu P, Li M, Xu X, Shi Y, Cheng Y, Wei S, Zheng Y, Xi T, Cai H, Liu Z. Bioinspired Anchoring Agnps Onto Micro-Nanoporous Tio2 Orthopedic Coatings: Trap-Killing Of Bacteria, Surface-Regulated Osteoblast Functions And Host Responses. *Biomaterials*. 2016 Jan 1;75:203-22.
- [25] Kose N, Çaylak R, Pekşen C, Kiremitçi A, Burukoglu D, Koparal S, Doğan A. Silver Ion Doped Ceramic Nano-Powder Coated Nails Prevent Infection In Open Fractures: In Vivo Study. *Injury*. 2016 Feb 1;47(2):320-4.
- [26] Liao J, Han R, Wu Y, Qian Z. Review Of A New Bone Tumor Therapy Strategy Based On Bifunctional Biomaterials. *Bone Research*. 2021 Mar 16;9(1):18.
- [27] Hsu Wk, Goldstein Cl, Shamji Mf, Cho Sk, Arnold Pm, Fehlings Mg, Mroz Te. Novel Osteobiologics And Biomaterials In The Treatment Of Spinal Disorders. *Neurosurgery*. 2017 Mar 1;80(3s):S100-7.

- [28] Ganau M, Holly Lt, Mizuno J, Fehlings Mg. Future Directions And New Technologies For The Management Of Degenerative Cervical Myelopathy. *Neurosurgery Clinics*. 2018 Jan 1;29(1):185-93.
- [29] Tran Pa, Sarin L, Hurt Rh, Webster Tj. Differential Effects Of Nanoselenium Doping On Healthy And Cancerous Osteoblasts In Coculture On Titanium. *International Journal Of Nanomedicine*. 2010 May 13:351-8.
- [30] Serra G, Morais L, Elias Cn, Semenova Ip, Valiev R, Salimgareeva G, Pithon M, Lacerda R. Nanostructured Severe Plastic Deformation Processed Titanium For Orthodontic Mini-Implants. *Materials Science And Engineering: C*. 2013 Oct 1;33(7):4197-202.
- [31] Puértolas Ja, Kurtz Sm. Evaluation Of Carbon Nanotubes And Graphene As Reinforcements For Uhmwpe-Based Composites In Arthroplastic Applications: A Review. *Journal Of The Mechanical Behavior Of Biomedical Materials*. 2014 Nov 1;39:129-45.
- [32] Garimella R, Eltorai Ae. Nanotechnology In Orthopedics. *Journal Of Orthopaedics*. 2017 Mar 1;14(1):30-3.
- [33] Lin Y, Taylor S, Li H, Fernando Ks, Qu L, Wang W, Gu L, Zhou B, Sun Yp. Advances Toward Bioapplications Of Carbon Nanotubes. *Journal Of Materials Chemistry*. 2004;14(4):527-41.
- [34] Singh K, Kim Kc. Early Detection Techniques For Osteoporosis. *Intech Open Access Publisher*; 2012 Feb 24.