

Background

- Osteoporosis and Fragility Fractures
- According to *Archives of Osteoporosis*, fragility fractures secondary to osteoporosis are highly prevalent, with about 9 million cases yearly worldwide
- The NHANES estimated 13.9 million people, above the age of 50, to suffer from Osteoporosis in 2020.
- Requirements for Ideal Solution
- Biodegradability
- Biocompatibility
- Mechanical strength

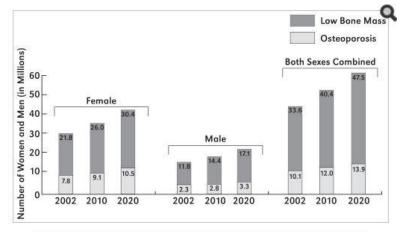
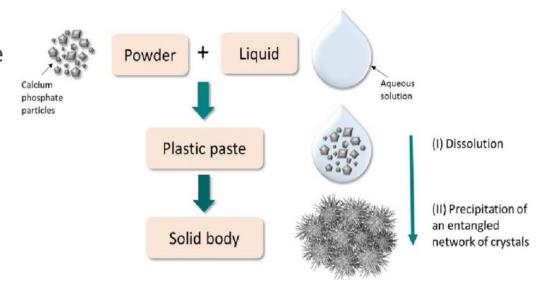


Figure 1: Projected Prevalence of Osteoporosis and/or Low Bone Mass of the Hip in Women, Men, and Both Sexes, 50 Years of Age or Older, *Bone Health and Osteoporosis: A Report of the Surgeon General.*

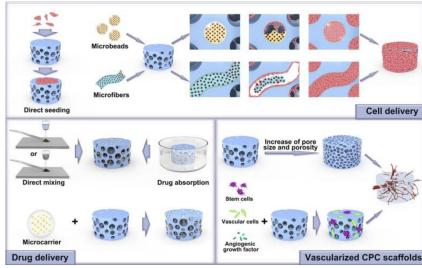
Proposed Solution

- What are calcium phosphate cements?
- Advantages of injectability
- Biocompatibility, bioactivity, biodegradability



Proposed Solution (cont)

- Osteoinductive properties
- Effective scaffold for bone tissue growth
- Versatility
 - Various additives can be introduced to improve certain properties of CPCs
 - Can be used in conjunction with larger fixation implant as potential solution for large fractures



Versatility of CPCs as a bioreactive therapy (*Eliaz*, et al., 2017).

Models for Testing

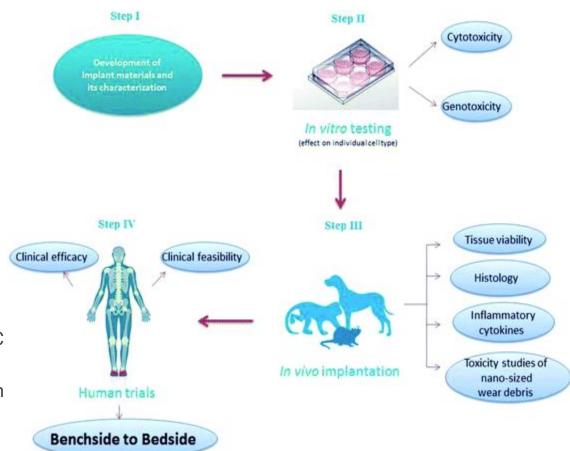
After developing the proposed CPC implant, the following stages of testing are required:

1) In vitro testing

 Test multiple cell lines to evaluate CPC's toxicity, debris, oxidative stress, etc on bone osteoblasts (ISO 10993-5:2009)

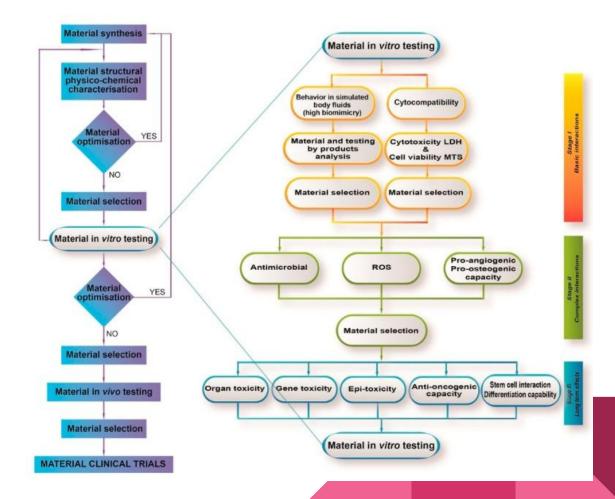
2) In vivo testing

- a) Use rat model to determine CPC effect in dermal implant
- b) Use canine model to test CPC in bone implant



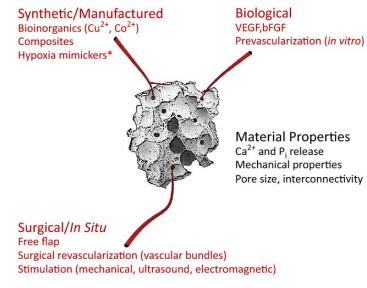
FDA Guidelines

- Developing any biomaterial that adheres to bone is prone to risk
- FDA has rigorous protocols for development and research



Limitations

- Mechanical properties- CPCs have are brittle, low Young's modulus.
 - Brittle issue because of biomechanicsjoints incur more stress than other parts of the body.
 - Mitigate using metal to bear the load
- Pre-vascularization necessary or cells die of hypoxia. Natural angiogenesis isn't fast enough.
 - Can use stem cells, but controversial.
 - Possibly mitigate with some localized drug to speed up angiogenesis?



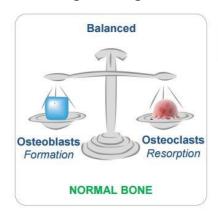
Trends in Biotechnology

le Ontione To Address Angiogenesis I leing Cap Rone-Graft Substitutes V



Future Directions

- Underlying Issue: Excessive bone resorption
 - Osteoclast inhibitor drugs
 - Bisphosphonates
 - Osteogenesis inducing drugs
 - Teriparatide and Denosumab
 - Drug Eluting device ensures proper dosing





References

Thrivikraman, G., Madras, G., & Basu, B. (2014, March 5). In vitro / In vivo assessment and mechanisms of toxicity of bioceramic materials and its wear particulates. Retrieved April 28, 2020, from https://pubs.rsc.org/en/content/articlehtml/2014/ra/c3ra44483i

Ginebra, Maria-Pau & Espanol, Montserrat & Maazouz, Yassine & Bergez, Victor & Pastorino, David. (2018). Bioceramics and bone healing. EFORT Open Reviews. 3. 173-183. 10.1302/2058-5241.3.170056.