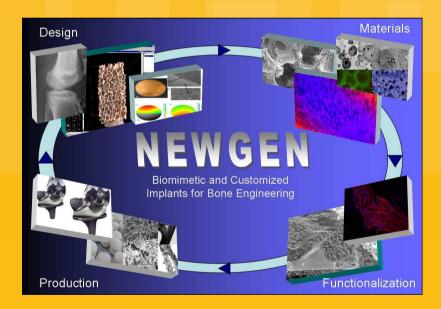


Additive manufacturing of bioceramic scaffolds:



State of the art

Eric Champion

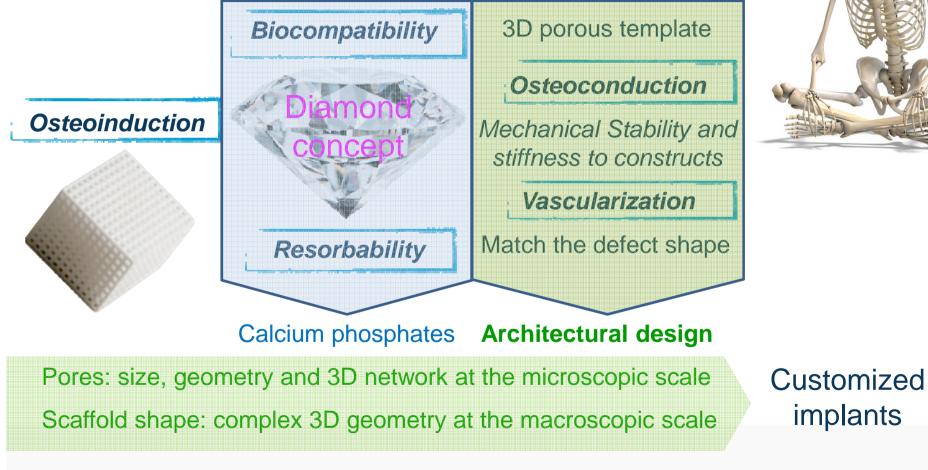




Background

Scaffolds for bone tissue engineering: several requirements

(PV Giannoudis, TA Einhorn, G Schmidmaier, D Marsh. The diamond concept - open questions. Injury, 2008) (JR Jones, LL Hench. Regeneration of trabecular bone using porous ceramics. Current Opinion Solid State Mater Sci, 2003)







Background

Additive Manufacturing

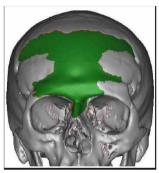
- Technologies implemented to produce prototypes or parts, within a short time, via an automated process.
- CAD/CAM using (generally) layer-by layer fabrication procedure.
- Initially set up for polymers and metals
- Some have been adapted to the shaping of 3D ceramic parts

Versatile technologies of particular interest in the biomedical field

Example. HA cranio-facial implant

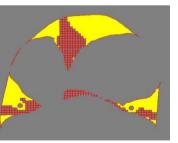






CAD of implant

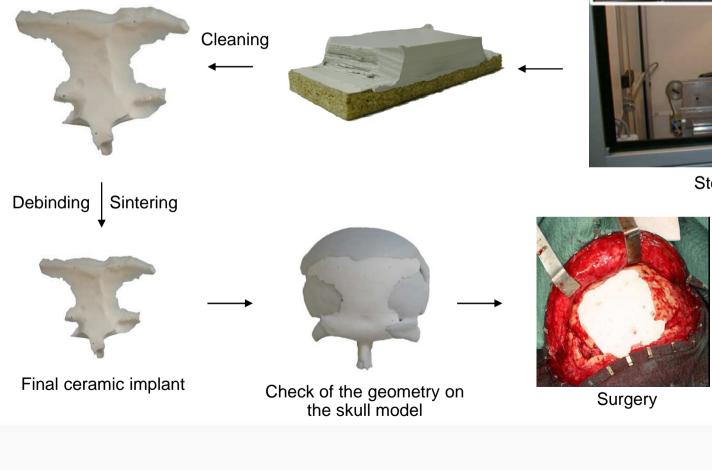




CAD in STL format (including shrinkages & strains) 2D Slicing

Background

Additive Manufacturing





Stereolithography



3D AM techniques	Tolerance	Advantages	Limitations
Material extrusion	1 mm	 Ease of support removal Good mechanical properties No material waste 	- Precision limited by the filament diameter
Selective laser sintering	0.2 to 0.5 mm	 High production rates, low cost Complex designs Good surface finishing 	 High roughness of the surface Poor mechanical properties Limited to materials which absorb IR light
Binder jetting	0.05 to 0.1 mm	 Wide variety of materials Simple technology 	 High roughness of the surface Expensive technology Poor mechanical properties Use of toxic organic binders
Vat photopolymerization	0.01 to 0.1 mm	 Complex designs Good surface finishing Good mechanical properties High accuracy 	 Expensive photosensitive resins Cleaning step necessary Control of the vertical accuracy

(T Chartier, C Dupas, M Lasgorceix, J Brie, E Champion, N Delhote, C Chaput. Review: Additive manufacturing to produce 3D ceramic parts. J Ceram Sci Tech, 2015)



Type of feedstock	Habitus	Rheological and physical properties	Notes about processability	Technology	Notes about composition
Powder Particle size > 20 μm >20 μm 80 μm>		Solid	Poor flowability	P-3DP P-SLS	Several possible binder systems,
			Good flowability	P-SLM	organic or inorganic based.
	<80 µm		Excellent flowability; high powder packing density		
Paste	Filament diameter 50 μm< <1000 μm	Viscosity: ^{107, 110, 146} 10–100 Pas @ 100 s ⁻¹ G' (eq) = 10^5 – 10^6 Pa Yield stress 10^2 – 10^3 Pa	Shear thinning	DIW Robocasting	Ceramic solid loading 35–55 vol%. Use of flocculated suspensions or of polymeric binders
	100 μm< <1000 μm			FDM	Typical ratio ceramic/polymer = 60/40 vol
10 μr <50 μ 10 μm <50 μ 50 μm	Layer thickness 10 μm< <50 μm	Viscosity: ^{11, 64} 100 mPa·s–110 Pa·s @ 100 s ⁻¹	Shear thinning, suitable viscosity highly depends on the recoating system	SL	Typical composition includes: monomer, photoinitiator, dispersants Ceramic solid loading 40–60 vol%
	10 μm< <50 μm	Viscosity: ^{97, 98, 147, 148} 5–15 mPa·s @ 1000 s ⁻¹ Surface energy: 20–70 mN/m	Shear thinning	DIP	Solids loading 2–30 vol%
	50 μm< <200 μm	Viscosity:1 mPa·s-1 Pa·s	Shear thinning	S-3DP S-SLS	Water basedCeramic solid loading 30–50 vol%

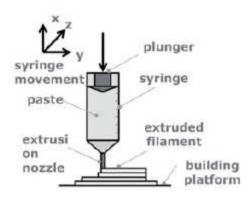
(A Zocca, P Colombo, C Gomes, J Günster. Additive manufacturing of ceramics: issues, potentialities and opportunities. J Am Ceram Soc, 2015)



Material Extrusion

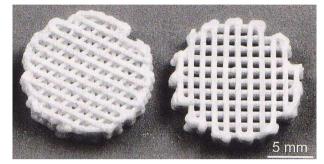
(Robotic dispensing, fused deposition modeling, 3D-printting, multiphase jet solidification)

Material (liquid or paste) is selectively extruded through a nozzle and deposited as rods

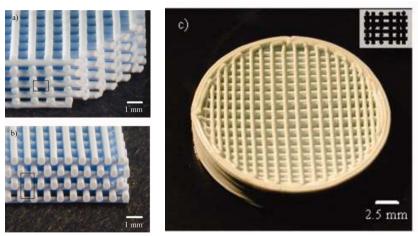


(A Zocca et al. J Am Ceram Soc, 2015)

Examples (sintered parts after shaping)



HA scaffolds with different pore geometries (U Deisinger *et al.* Key Engng Mat, 2008)



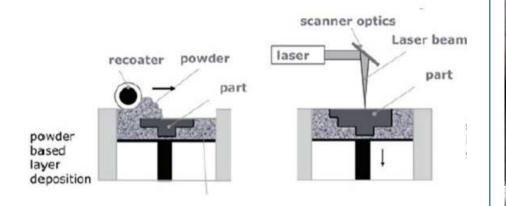
HA scaffold with multiscale porosity LOW accuracy (JG Dellinger *et al.* J Biomed Mater Res A, 2007)



Selective Laser Sintering

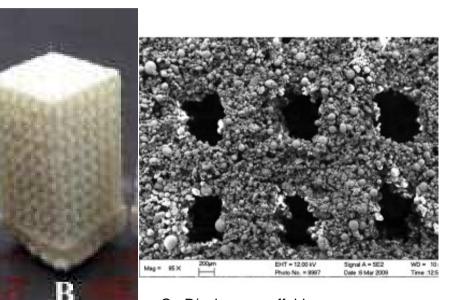
(Selective laser sintering, selective laser melting, electron beam metling)

Thermal energy (laser or electron beam) selectively sinters or melts scanned regions of a powder bed mixed with a binder



(A Zocca et al. J Am Ceram Soc, 2015)

Example



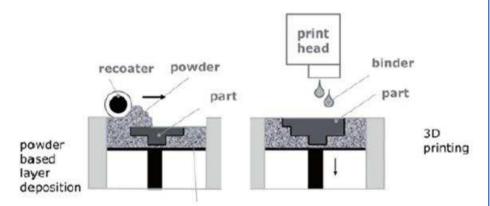
Ca-P/polymer scaffold. (B Duan *et al.* Acta Biomater, 2010)

Low accuracy and mechanical properties

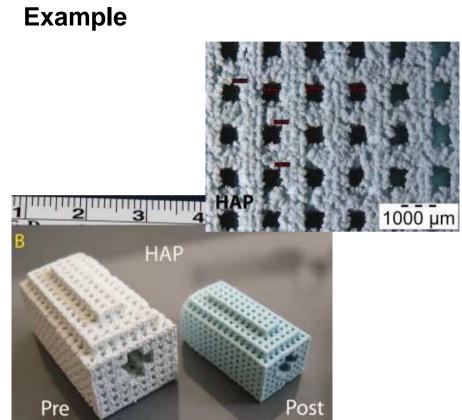


Binder Jetting

A liquid bonding agent is selectively deposited to consolidate a powder bed



(A Zocca et al. J Am Ceram Soc, 2015)



HA scaffold after shaping (Pre) and after sintering (Post). (PH Warnke *et al.* J Biomed Mater Res B Appl Biomater, 2010)

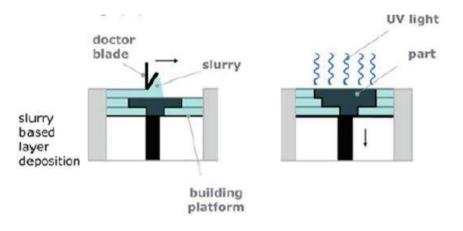
Low mechanical properties



Vat Photopolymerization

(stereolithography, microstereolithography)

A reactive suspension is selectively cured by light-activated polymerization



⁽A Zocca et al. J Am Ceram Soc, 2015)

Examples Image: Constraint of the second sec

HA scaffolds (bone substitute & occular implant)

- Number of commercially available products
- On-demand large implants with complex shape

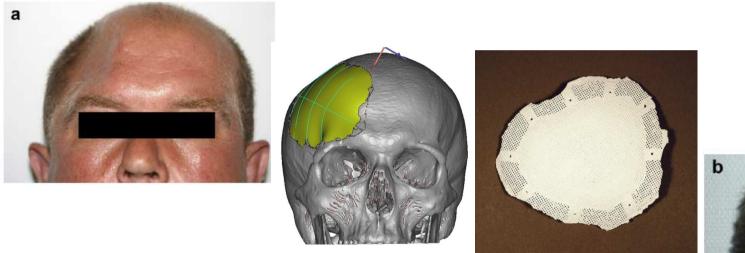
Good accuracy and mechanical properties



Vat photopolymerization (Stereolithography)

Example: On-demand cranio-facial implants made in hydroxyapatite





Efficiency of SLA to produce large complex ceramic implants or scaffolds but...



(J Brie, T Chartier, C Chaput, C Delage, B Pradeau, F Caire, MP Boncoeur, JJ Moreau. A new custom made bioceramic implant for the repair of large and complex craniofacial bone defects. J Cranio-Maxillofac Surg, 2013)

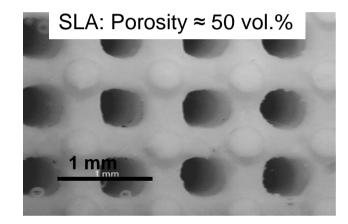


Vat photopolymerization (Stereolithography)

Enhance new tissue ingrowth

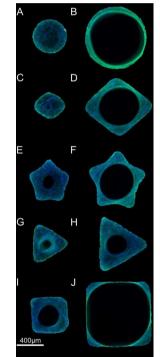
Increase the dimensionnal resolution: microstereolithography

(SLA \approx 100 µm - µSLA \approx 10 µm)



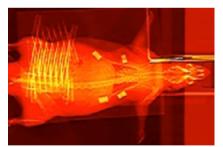
Highly porous scaffolds (Porosity > 70%) & Adjusted geometry and network of pores

In vitro cell proliferation





Ex-ovo vascularization



In vivo integration

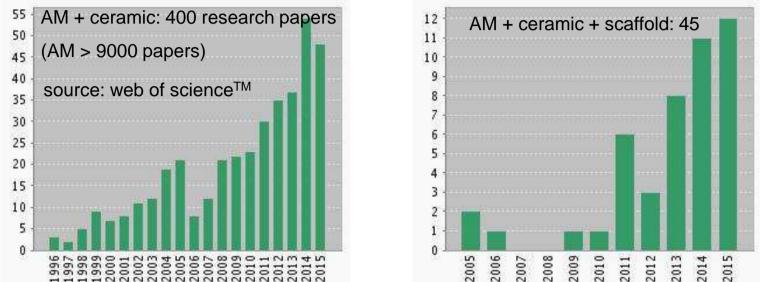
M Lasgorceix, PhD, SPCTS Limoges collaboration Dr U Rudrich & Dr A Magnaudeix





Concluding remarks

> AM of ceramic scaffolds: very young technologies



Few of them have reached the clinic and led to commercially available products

Processing requirements are still very challenging (feedstock, sintering...)

Promising technologies to the reliable production of bioceramics:

On-demand large implants with accurate dimensions Scaffolds with controlled pore size and geometries Multiscale control of architectures

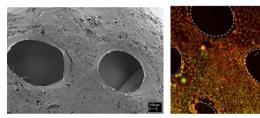


Concluding remarks

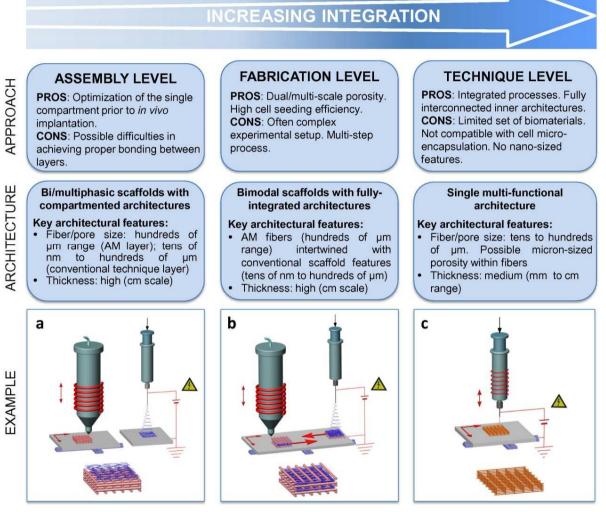
Further developments

Combination of several AM technologies

Association with cells seeding using biofabrication technologies (bioprinting, robocasting..)



(T Zehnder, B Sarker, AR Boccaccini, R Detsch. Evaluation of an alginate–gelatine crosslinked hydrogel for bioplotting. Biofabrication, 2015)



(SM Giannitelli, P Mozetic, M Trombetta, A Rainer. Combined additive manufacturing approaches in tissue engineering. Acta Biomater, 2015)



SPCTS

Thanks for your attention

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Joël Brie, Thierry Chartier, Marie Lasgorceix

