

組織工程用多孔性生醫玻璃陶瓷支架之研究

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摘要

由於外傷、疾病或者手術所造成組織的損壞經常需要進行骨移植，而治療的方式有自體移植、異體移植與異種移植等。人工支架材料可促使骨再生及支撐新生骨，而3D支架材料需有高孔隙度、開放性結構並且能使植入材有適當的血管增生，此外支架材料也需能使養份與排泄物流通。作為引導骨再生的支架用材料中，最常被研究探討的有氫氧基磷灰石、三鈣磷酸鹽、生物活性玻璃和含有磷灰石與矽灰石的玻璃陶瓷等。本實驗是發展以稻殼作為成孔劑添加到生物活性玻璃陶瓷45S5R中製作多孔性支架材料。實驗結果顯示，玻璃分別經過450 °C 與1050 °C、1小時的結晶熱處理後，主要晶相種類以XRD測定結果為：磷灰石、Na₂Ca₂Si₃O₉兩種結晶相。以添加三種不同含量及粒徑大小的稻殼經過1050 °C、1小時燒結後，分別利用SEM、阿基米德原理量測孔隙大小、開放型孔隙度。以添加SA75P3的稻殼成孔劑，其大孔隙分別為長軸：600 ± 15 μm 短軸：65 ± 25 μm、開放型孔隙度則為47.2 ± 2.7%。由型態組織觀察的結果顯示，SA75P3試片的孔隙大小分佈、孔隙之互相連通性與表面粗糙度方面都適合作為骨組織工程用的支架材料。

關鍵詞：玻璃陶瓷；支架材料；多孔性

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- [1] Wiss DA. Fractures. In: Capers CM, Williams CB, editors. New York: Lippincott-Raven Publishers 1998.p.240.
- [2] Stanford CM. J Am Dent Assoc 2005;136:1092-1100.
- [3] Melba N, Sergio V, Salvador M, Maria GP. New macroporous calcium phosphate glass ceramic for guided bone regeneration. Biomaterials, 2004;25(18):4233-4241.
- [4] Ishizaki K, Komarneni S, Nanko M. Porous Materials: Process Technology and Applications. Dordrecht: Kluwer Academic Publishers 1998.p.181-198.
- [5] Kingery WD, Bowen HK, Uhlmann DR. In: introduction to Ceramic. 2nd ed. Wiley, New York 1976.p.368.
- [6] Park JB, Lakers RS. Biomaterials: An Introduction. 2nd Plenum press, New York. 1992.
- [7] Hench LL. Bioceramics: From concept to clinic. J Am Ceram Soc 1991;74:1487-1570.
- [8] Lin FH, Lin CC, Liu HC, Huang YY, Wang YY. Sintered porous-bioglass & hydroxyapatite as bone substitute. Biomaterials 1994;15(13):1087-1098.
- [9] Hench LL, Splinter RJ, Allen WC, Greenlee TK. Bonding mechanism at the interface of ceramic prosthetic materials. J Biomed Mater Res 1971;2:117-141.
- [10] Broemer H, Deutscher K, Blencke B, Pfeil E, Strunz V. Properties of the bioactive implant material ‘Ceravital’. J Sci Ceram 1997;9:219-225.
- [11] Abe Y, Hosoe M, Kasuga T. High strength Ca(PO₃)₂ glass-ceramics prepared by unidirectional crystallization. J Am Ceram Soc 1982;65:189-190.
- [12] Vogel W. Chemistry of glass. J Am Ceram Soc 1985.p.251.
- [13] Kokubo T, Ito S, Sakka S, Yamamoto T. Formation of a high-strength bioactive glass-ceramic in the system MgO-CaO-SiO₂-P₂O₅. J Mater Sci 1986;21:536-540.
- [14] Lin FH, Hon MH. A study on the bioglass ceramics in the NaO-CaO-SiO₂-P₂O₅ system. J Mater Sci 1989;23:4295-4299.
- [15] Feenstra L, de Groot K. Medical use of calcium phosphate ceramics. In: Bioceramics of Calcium Phosphate, CRC Press, 1983.p.131-141.
- [16] Peltier LE. The use of plaster of paris to fill defect in bone. Clin Orthop 1961;21:10-31.
- [17] Graves GA, Hentrich RL, Stain HG, Baijapai PK. Resorbable ceramic implants in bioceramic. Engineering in Medicine. New York: Interscience Publisher, 1972.p.91-115.
- [18] Jarcho M, Salsbury RL, Thomas MB. Synthetic and fabrication of B-TCP ceramic for potential prosthesis application. J Mater Sci 1979;14:142-150.
- [19] Chohayeb AA, Chow LC, Tsaknins PJ. Evaluation of calcium phosphate as a root canal sealer-filler material. J Endodont 1987;13:384-387.
- [20] Hong YC, Wang JT, Hong CY, Brown WE, Chow LC. The periapical tissue reactions to a calcium phosphate cement in the teeth of he monkeys. J Biomed Mater Res 1991;25:485-498.
- [21] Miyamoto Y, Toh T, Ishikawa K, Yuasa T, Nagayama M, Suzuki K. Effect of added NaHCO₃ on the basic properties of apatite cement. J Biomed Mater Res 2001;2:147-151.
- [22] Knaack D, Goad MEP, Aiolova M, Rey C, Tofiqhi A, Chakravarthy P, Lee DD. Resorbable calcium phosphate bone substitute. J Biomed Mater Res 1998;43:399-409.
- [23] Zhou J, Zhang X, Chen J, Zeng S, de Groot K. High temperature characteristics of synthetic hydroxyapatite. J Mat Sci:Mat in Med 1993;4:83-85.
- [24] Van Mullen PJ, de Wijn JR, Vaandrager JM. Porous acrylic cement: Evaluation of a novel implant material. Ann Plast Surg Dec 1988;21(6):576-582.
- [25] Park JB. J Biomater Sci and Engineering, Plenum Press:New York and London, 1985.

- [26] 許鎮龍, 潘志明, 彭康祐, 江國威. 稻殼及造紙廠廢水污泥運用於鍋爐燃料之可行性探討. 聯合學報, 2002.p. 89-194.
- [27] 台灣省政府農林廳. 民國八十六年版台灣農業年報, 1997.
- [28] 楊紹榮. 農業廢棄物處理與再利用. 台南區農業改良場, 2006.
- [29] 謝惠子. 黃金稻田的秘密-雲林縣稻殼氣化發電廠. 能源報導, 2004.p. 8-10.
- [30] 王岱淇. 玉米穗軸與稻殼再生質燃燒爐之燃燒分析. 臺灣大學農業機械工程研究所碩士論文, 台北, 1993.
- [31] 吳中興, 李志萍. 使用旋轉窯焚化爐焚化稻殼之數值模擬分析. 農業機械學刊第13卷第1期, 台北, 2004.
- [32] Fino D, Russo N, Saracco G.. A multifunctional filterfor the simultaneous removal of fly-ash and NOx from incinerator flue gases. Chem Eng Sci 2004;59(22-23):5329-5336.
- [33] Anappara AA, Rajeshkumar S, Mukundan P, Warrier PRS. Mpedance spectroscopic studies of sol-gel derived subcritically dried silica aerogels, Acta Mater 2004;52(2):369-375.
- [34] Haugen H, Will J, Kohler A, Hopfner U, Algner J, Wintermantel E. Ceramic TiO₂-foams: characterization of a potential scaffold. J Eur Ceram Soc 2004;24(4):661-668.
- [35] Schwartzwalder K, Somers H, Somers AV. Method of making porous ceramic articles. USA Patent No.3090094, 1963.
- [36] 朱新文, 江東亮. 有機泡沫浸漬工藝一種經濟實用得多孔陶瓷製作工藝. 砂酸鹽通報, 2000.p. 45-51.
- [37] Falamaki C, Naimi M, Aghaie A. Dual behavior of CaCO₃ as a porosifier and sintering aid in the manufacture of alumina membrane/catalyst upports. J Eur Ceram Soc 2004;24(10-11):3195-3201.
- [38] Diaz A, Hampshire S. Characterization of porous silicon nitride materials produced with starch. J Eur Ceram Soc 2004;24(2):413-419.
- [39] 資文華, 孫俊賽, 黃明華. 溶膠-凝膠法製作多孔陶瓷的研究發展. 中國陶瓷, 2003,39(4):14-18.
- [40] Zu LJ, Luo SJ. Study on the powder mixing and semi-solid extrusion forming process of SiCp/2024 Al composites. J Mater Proc Tech 2001;114(3):189-193.
- [41] Kim T, Goto T, Lee B. Microstructure control and mechanical properties of fibrous Al₂O₃/ZrO₂ composites fabricated by extrusion process. Scripta Mater 2005;52(8):725-729.
- [42] Qiao GJ, Ma R, Jin ZH. Microstructure transmissibility in preparing SiC ceramics from natural wood. J Mater Proc Tech 2002;120(1-3):107-110.
- [43] Qian JM, Wang JP, Qiao GJ, Jin ZH. Preparation of porous SiC ceramic with a woodlike microstructure by sol-gel and carbothermal reduction processing. J Eur Ceram Soc 2004;24(10-11):3251-3259.
- [44] Luhlech H, Dias J, Nickel H. The coat-mix procedure using carbon fillers. Carbon 1997;35:95-102.
- [45] Simwonis D, Thulen H, Dias FJ, Naoumidis A, Stover D. Properties of Ni/YSZ porous cermets for SOFC by tape casting and coat-mix process. J Mater Proc Tech 1999;92(93):107-111.
- [46] Middleton H, Diethelm S, Ihinger R. Co-casting and co-sintering of porous MgO support plates with thin dense perovskite layers of LaSrFeCoO₃. J Eur Ceram Soc 2004;24(6):1083-1086.
- [47] Fukasawa T, Deng ZY, Ando M. Pore structure of porous ceramics synthesized from water-based slurry by freeze-dry process. J Mater Sci 2001;36:2523-2527.
- [48] 徐仁輝. 粉末冶金概論. 新文京開發出版有限公司, 2002.
- [49] 汪建民. 粉末冶金技術手冊. 中華民國冶金協會, 1999.P.200.
- [50] 伍祖聰, 黃錦鐘. 粉末冶金. 高立圖書有限公司, 1996.P.193.
- [51] 汪建民. 粉末冶金技術手冊. 中華民國冶金協會, 1999.P.202.
- [52] 伍祖聰, 黃錦鐘. 粉末冶金. 高立圖書有限公司, 1996. P.176.
- [53] German RM. Powder Metallurgy Science, Metal Powder Industries Federation Princeton. New Jersey 08540-6692 U.S.A. 1994.
- [54] Sepulveda P, Jones JR, Hench LL. Bioactive sol-gel foams for tissue repair. J Biomed Mater Res A 2002;59(2):340-348.
- [55] Jones JR, Hench LL. Regeneration of tragecular bone using porous ceramics. Curr Opin Solid State Mater Sci 2003;7:301-307.
- [56] Clupper DC, Hench LL. Crystallization kinetics of tape cast bioactive glass 45S5. J Non-Crys Solid 2003;318:43-48.
- [57] Jones JR, Hench LL. Effect of surfactant concentration and composition on the structure and properties of so-gel-derived bioactive glass foam scaffolds for tissue engineering. J Mater Sci 2003;38:1-8.
- [58] Saravanapavan P, Hench LL. Mesoporous calcium silicate glasses: I synthesis. J Non-Cryst Sol 2003;318:1-13.
- [59] Jones JR, Hench LL. Factors affecting the structure and properties of bioactive foam scaffolds for tissue engineering. J Biomed Mater Res B: Apply Biomater 2004;68B:36-44.
- [60] Clupper DC, Hench LL, Mecholsky JJ. Strength and toughness of tape cast bioactive glass 45S5 following heat treatment. J Eur Ceram Soc 2004;24:2929-2934.
- [61] Jones JR, Ehrenfried LM, Hench LL. Optimising bioactive glass scaffolds for bone tissue engineering. Biomaterials 2006;27:964-973.
- [62] 王明光, 王敏昭. 實用儀器分析. 合記圖書出版社, 2003.
- [63] Ozguur EN, Cuuneyt TA. Manufacture of Macroporous Calcium Hydroxyapatite Bioceramics. J Eur Ceram Soc 1999;19:2569-2572.
- [64] Chen QZ, Thompson LD, Boccaccini AR. 45S5 BioglassR-derived glass-ceramic scaffolds for bone tissue engineering. Biomaterials

2006;27:2414-2425.

- [65] 葉郁仁, 林鴻儒, 郭俊榕, 楊俊佑, 吳侑峻. 應用於骨組織工程Alginate/HAP多孔海綿體之製備及活體外與活體內相關性質之研究. 中華民國生物醫學工程學會, 2002.
- [66] Mears DC. Metals in medicine and surgery. *Int Metal Rev* 1997;22:119-153.
- [67] Tampieri A, Celotti G, Sprio S, Delcogliano A, Franzese S. Porosity-graded hydroxyapatite ceramics to replace natural bone. *Biomaterials* 2001;22:1365-1370.
- [68] Xin R, Leng Y, Chen J, Zhang Q. A comparative of calcium phosphate formation on bioceramics in vitro and in vivo. *Biomaterials* 2005;26:6477-6486.
- [69] Ding SJ, Wang CW, Chen CH, Chang HC. In vitro degradation behavior of porous calcium phosphates under diametral compression loading. *Ceram Int* 2005;31:691-696.
- [70] Yuan H, Bruijn JD, Zhang X, Blitterswijk CA, Groot K. Bone Induction by Porous Glass Ceramic Made from BioglassR (45S5). *J Biomed Mater Res* 2001;58:270-276.
- [71] Kokubo T, Ito S, Huang ZT, Hayashi T, Sakka S, Kitsugi T, Yamamoto T. Ca, P-rich layer formed on high-strength bioactive glass-ceramic A-W. *J Biomed Mater Res* 1990;24:331-341.
- [72] Chen QZ, Rezwan K, Francon V, Armitage D, Nazhat SN, Jones FH, Boccaccini AR. Surface functionalization of BioglassR -derived porous scaffolds. *J Acta Biomater* 2007;3:551-562.
- [73] Clupper DC, Mecholsky Jr JJ, Latorre GP, Greenspan DC. Bioactivity of tape cast and sintered bioactive glass-ceramic in simulated body fluid. *Biomaterials* 2002;23:2599-2606.
- [74] Chen QZ, Thompson ID, Boccaccini AR. 45S5 BioglassR derived glass-ceramic scaffolds for bone tissue engineering. *Biomaterials* 2006;27:2414-2425.
- [75] Lefebvre L, Chevalier J, Grmillard L, Zenati R, Thollet G, Bernache-Assolant D, Govin A. Structural transformations of bioactive glass 45S5 with thermal treatments. *J Acta Mater* 2007;55:3305-3313.
- [76] Wu SC, Wang CL, Hon MH. Effects of Ca/P ratio on the crystallization of MgO-CaO-Al₂O₃-SiO₂-P₂O₅ glass-ceramics. *J Ceram Soc of Japan* 1995;103(2):99-103.
- [77] German RM. Particle packing characteristics. NJ: Metal Powder Industries Federation 1989;404-407.
- [78] Martin CL, Bouvard D. Isostatic compaction of bimodal powder mixtures and composites. *Int J Mech Sci* 2004;24:907-927.
- [79] McGeary RK. Mechanical packing of spherical particles. *J Am Ceram Soc* 1961;44:513-522.
- [80] 汪建民, 朱秋龍. 粉末冶金. 中華民國粉末冶金協會, 1999.
- [81] Cai Y, Zhou L. Effect of thermal treatment on the microstructure and mechanical properties of gel-derived bioglasses. *J Mater Chem and Phys* 2005;94:283-287.
- [82] Jones JR, Ehrenfried LM, Hench LL. Optimising bioactive glass scaffolds for bone tissue engineering. *Biomaterials* 2006;27:964-973.
- [83] Gong W, Abdelouas A, Lutze W. Porous bioactive glass and glass-ceramics made by reaction sintering under pressure. *J Biomed Mater Res* 2001;54:320-327.
- [84] Xynos ID, Hukkanen MV, Batten JJ, Buttery LD, Hench LL, Polak JM. Bioglass 45S5 stimulates osteoblast turnover and enhances bone formation In vitro: implications and applications for bone tissue engineering. *Calcified Tissue International* 2000;67:321-329.
- [85] El-Ghannam AR. Advanced bioceramic composite for bone tissue engineering: design principles and structure-bioactivity relationship. *J Biomed Mater Res* 2004;69:490-501.
- [86] Yuan H, de Bruijn JD, Zhang X, van Blitterswijk CA, de Groot K. Bone Induction by Porous Glass Ceramic Made from BioglassR (45S5). *J Biomed Mater Res* 2001;58:270-276.
- [87] Hench LL, Paschall HA. Direct chemical bond of bioactive glass ceramic materials to bone and muscle. *J Biomed Mater Res Symposium* 1973;7:25-42.
- [88] Hench LL, Paschall HA. Histochemical response at a biomaterial 's interface. *J Biomed Mater Res Symp* 1974;55:49-64.
- [89] Wilson J, Piggot GH, Schoen FJ, Hench LL. Toxicology and Biocompatibility of Bioglass. *J Biomed Mater Res* 1981;15:805-810.
- [90] Hench LL, West J. The sol-gel process. *Chem Rev* 1990;90:33-72.
- [91] Hench LL, West J. Biological applications of bioactive glasses. *Life Chemistry Reports* 1996.p. 187.
- [92] Hench LL. Sol-gel materials for bioceramic applications. *Curr Opin Solid State Mater Sci* 1997;2:604-610.
- [93] Oonishi H, Kutrshitan S, Yasukawa E, Iwaki H, Hench LL, Wilson J, Tsuji E, Sugihara T. Particulate bioglass compared with hydroxyapatite as a bone graft substitute. *Clin Orthop Relat res* 334:316-325.
- [94] Hench LL. Bioceramics. *J Am Ceram Soc* 1998;81:1705-1728.
- [95] Hench LL. Bioactive materials: the potential for tissue regeneration. *J Biomed Mater Res* 1998;41:511-518.
- [96] Coleman NJ, Hench LL. A gel-derived mesoporous silica references materials for surface analysis by gas sorption, part 1-textural features. *Ceram Int* 2000;26:171-178.