



7~13. 1311标记CD133单链抗体对人肝癌CD133 +HepG2干细胞的抑制作用[J].侯妍利,陈兴月,段丽群,唐敏,康强强,舒锦,李少林(1)

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摘要:

目的: 研究1311标记CD133单链抗体(single chain variable fragment, ScFv)在体外对人肝癌CD133 + HepG2干细胞的抑制作用, 流式细胞术检测分选前后HepG2细胞的CD133表达率, 克隆形成实验及体内成瘤实验验证 CD133 + HepG2细胞133 ScFv并测定标记率、比活度、放射性浓度。将分选出的CD133 + HepG2细胞分为1311-CD133抗体治疗组、1311治疗组、33抗体治疗组, MTT法检测各组中对CD133 + HepG2细胞生长抑制的最适剂量和不同药物在12、48、72 h三个时间点对CD133用, 流式细胞术检测各组细胞周期的变化。结果: 分选的HepG2细胞的CD133表达率显著高于未分选细胞[(97.71±1.13)% vs (1.52±0.78)% P=0.0001]。CD133 + HepG2细胞相对于CD133 - HepG2细胞具有更强的体外成球、克隆形成能力[(45.03±1.35)% vs (7.4±0.54)% P<0.001]和体内成瘤能力。1311-CD133 ScFv的标记率为88.92%, 放射化学纯度为98.63%。当1抗体为1 μg/100 μl时, 对CD133 + HepG2细胞的抑制率最高, 达(89.58±0.74)%; 在此剂量下1311-CD133 ScFv生长抑制率显著高于其余各实验组, 且呈时间依赖性。1311-CD133 ScFv治疗组 G 0/G 1期细胞比例为(27.50±1.12)%, (P<0.05)。结论: 成功制备的1311-CD133 ScFv在体外能有效抑制人肝癌CD133 + HepG2干细胞的生长。

关键词: [肝癌干细胞](#) [CD133](#) [1311](#) [单克隆抗体](#) [单链抗体](#)

Inhibitory effect of 1311-labeled anti-CD133 ScFv on CD133 + human hepatocellular carcinoma cells

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Abstract:

Objective: To study the inhibitory effect of the anti-CD133 single chain variable fragment (ScFv) labeled with <sup>131</sup>I on CD133-positive human hepatocellular carcinoma stem cells (CSCs) sorted from human hepatocellular liver carcinoma HepG2 cells in vitro. Methods: CD133-positive CSCs were isolated from HepG2 cells through magnetic-activated cell sorting (MACS). CD133 expression in both sorted and unsorted cells was analyzed by flow cytometry (FCM). The property of CD133 + CSCs was validated by sphere-forming assay in vitro and tumor formation assay in nude BALB/c mice in vivo. The monoclonal antibody CD133 was labeled with <sup>131</sup>I using the chloramines T method and the labeling rate, specific activity and radioactivity were evaluated. CD133 + CSCs were treated with CD133 ScFv, <sup>131</sup>I-CD133 ScFv, and <sup>131</sup>I + CD133 ScFv. At 12, 24 and 48 hours after treatment, cell proliferation was assessed by MTT assay and FCM respectively. Results: CD133 was detected in (97.71±1.13)% of the sorted CSCs but in only (1.52±0.78)% of unsorted HepG2 cells (P=0.0001). As compared with CD133 - HepG2 cells, CD133 + CSCs had a higher tumor sphere formation ability [(45.03±1.35)% vs (7.4±0.54)% P<0.001]. The <sup>131</sup>I-labeled CD133 ScFv showed a maximal CD133 + cell growth inhibition of (89.58±0.74)% (P<0.05) when <sup>131</sup>I was used at 3.7 MBq/100 μl and CD133 ScFv was used at 1 μg/100 μl, significantly higher than other groups. The proportion of G 0/G 1 phase arrest in cells treated with <sup>131</sup>I-CD133 ScFv was significantly reduced as compared with other groups (P<0.05). Conclusion: Radioisotope <sup>131</sup>I-labeled CD133 ScFv may effectively inhibit growth of CD133-positive human hepatocellular carcinoma cells in vitro.

Keywords: [liver cancer stem cell](#) [CD133](#) [1311](#) [monoclonal antibody](#) [single chain variable fragment\(ScFv\)](#)

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