

Enhanced in-vivo C13 spectroscopy using adiabatic INEPT sequences and custom-made RF coils

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Introduction: ¹³C spectroscopy offers several advantages compared to ¹H MRS, including increased spectral dispersion, and eliminates the need for water suppression, though it suffers from inherently low sensitivity. In addition, ¹³C spectroscopy is important from a physiological point of view because it offers the ability to measure metabolic fluxes such as those of the TCA cycle [1]. In this abstract, we report results obtained with a high-quality custom-made RF coil and optimized sequences including INEPT [2] (with square pulses) and adiabatic INEPT with BIR-4 pulses (BINEPT) [3, 4].

Materials and Methods: All experiments were performed on a 9.4 T horizontal bore (20 cm) animal scanner. We employed a high-quality custom-made RF probe, custom made by the third author, with an inner solenoid for ¹³C detection and an outer saddle for ¹H detection. To increase ¹³C signal we employed INEPT [2] and adiabatic INEPT with BIR-4 pulses [3, 4]. BIR-4 pulses of 400 μ s having maximum amplitude of 16 kHz and 4 kHz for ¹³C and ¹H, respectively, were synthesized on a Bruker Avance console. The intra and inter-pulse delays were set according to the ¹³C-¹H scalar coupling of acetate (130 Hz). Both BINEPT with direct ¹³C detection and double BINEPT for indirect ¹³C detection via ¹H were designed and tested. ¹H decoupling with adiabatic WALTZ-16 sequence was employed during acquisition.

Results: In phantoms our BINEPT sequence (Figure 1) resulted in increased sensitivity by a factor of 2.8 for ¹³C, when compared with direct excitation following a 90° pulse (Figures 2 and 3a). Similar results were obtained *in vivo* on a mouse (natural-abundance ¹³C spectra shown in Figure 3b). The optimized BINEPT sequence showed significant signal increase over INEPT and direct excitation (¹H decoupled). The BINEPT sequence also acted as a spectral editing sequence suppressing the carbons that do not have a direct proton attached, such as COOH (Figures 2b and 3b).

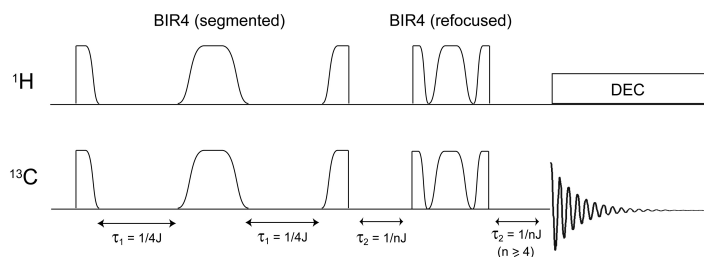


Figure 1. Adiabatic INEPT using segmented adiabatic BIR4 pulses. ¹³C signal is increased through polarization transfer via scalar coupling from ¹H.

Figure 2a. ¹³C spectra of [1,2-¹³C] labeled acetic acid: in red adiabatic INEPT, and in black direct excitation. An increase by a factor of 2.8 is obtained. Note that adiabatic INEPT acts also as a spectral editing sequence suppressing the carbons that do not have a direct proton attached, such as COOH.

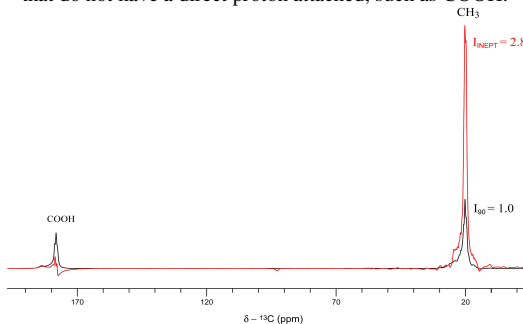


Figure 3a. Comparison of direct excitation ¹³C 90° pulse (¹H-decoupled), ¹³C INEPT and ¹³C BINEPT on a phantom containing [2-¹³C] labeled acetate. The BINEPT peak is 170% of the INEPT peak (the acetate peak has been set at 0 ppm).

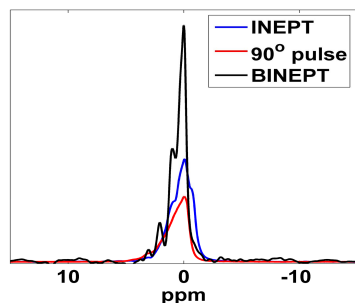


Figure 2b. ¹H spectra of [1,2-¹³C] labeled acetic acid: in red double adiabatic INEPT, and in black direct excitation. An efficiency of 30% is obtained compared to direct excitation. However, proton detection has superior sensitivity over the ¹³C from Figure 2a.

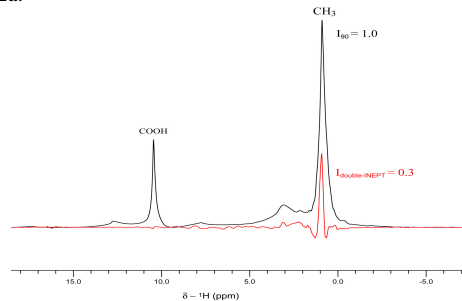
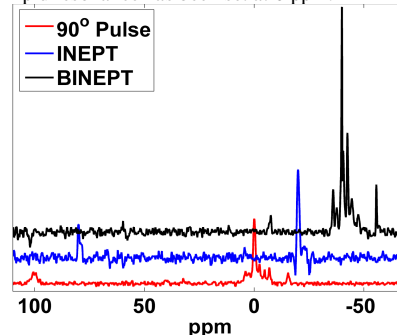


Figure 3b. *In vivo* ¹³C spectra of a mouse, collected with direct excitation ¹³C 90° pulse (¹H-decoupled), ¹³C INEPT and ¹³C BINEPT. The main BINEPT peak is 320% of the direct excitation peak (NEX=256, 10min total acquisition time). The lipid resonance has been set at 0 ppm.



Discussion: INEPT results in increased ¹³C signal through manipulation of the scalar coupling between ¹H and ¹³C to transfer polarization from ¹H to ¹³C. BINEPT (INEPT with adiabatic BIR-4 pulses) results in significant further ¹³C signal enhancement due to improvement of RF inhomogeneity, resulting in highly uniform spin excitation. Further improvements are possible, involving inverse detection of ¹³C through ¹H and/or extension to multidimensional heteronuclear experiments, useful for unambiguous assignment and quantification of *in vivo* spectra.

References:

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