

# Double-Quantum Filter Imaging of Sodium-23

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The method of double-quantum filter <sup>23</sup>Na NMR spectroscopy has been applied to <sup>23</sup>Na imaging. We have discriminated between <sup>23</sup>Na in the extreme narrowing condition (Na<sup>+</sup> in glycerol solution) and <sup>23</sup>Na in the slow motion condition (Na<sup>+</sup> in an albumin solution). The results suggest the possibility of correlation-time mapping of <sup>23</sup>Na.

## 1. Introduction

The method of double-quantum filter  $^{23}$ Na NMR spectroscopy is selective for the double-quantum coherences from the |1/2><-3/2|, |3/2><-1/2| rank 3 coherences. When  $\omega\tau_c$  is around or larger than 1 (the slow motion condition, where  $\omega$  is the Larmor frequency and  $\tau_c$  is the correlation time of  $^{23}$ Na), double-quantum coherence is evoked by using a double-quantum filter, When  $\omega\tau_c$  is much less than 1 (the extreme narrowing condition), double quantum coherence cannot be evoked $^{1-5}$ . The double-quantum filter has been applied in erythrocytes $^{6}$ , salivary gland $^{7}$ ) and heart $^{8}$ ), and the double-quantum coherences from the intracellular  $^{23}$ Na and  $^{39}$ K have been detected. In this study, we have applied this technique to  $^{23}$ Na imaging, and have tried to discriminate between  $^{23}$ Na in the extreme narrowing condition and  $^{23}$ Na in the slow motion condition. The results raise the possibility of correlation-time mapping of  $^{23}$ Na.

### 2. Materials and Methods

Sodium experiments were performed on a phantom consisting of two solutions: 1) an albumin solution (0.8 g albumin (Sigma A2153) and 1 ml 1 M NaCl solution) and 2) a glycerol solution (1 g glycerol and 1 ml 2 M NaCl solution). Each solution was placed in a methacrylate cuvette ( $1 \times 1$  cm square in cross section with a height of 1.5 or 1.2 cm). A Biospec 150/4.7 spectrometer (Oxford Research Systems, 4.7 T) was used with a home-built  $^{23}$ Na probe with a 20 mm internal

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diameter, 4-turn solenoidal transmitter/receiver coil of 2.0 mm copper wire. Two variable capacitors (Voltronics, 38pF) were used for tuning (53.0 MHz) and matching (50 $\Omega$ ).

### 3. Results and Discussion

Since  $\omega \tau_c$  of <sup>23</sup>Na in the albumin solution is larger than 1 (the slow motion condition) at 25°C, <sup>23</sup>Na in the albumin solution showed a signal from the double-quantum coherence. The spin-echo double-quantum filter, d-90°- $\tau$ /2-180°- $\tau$ /2-90°- $\delta$ -90°-acquire was applied with a 32-step phase cycle<sup>2,9)</sup>. The transverse relaxation rate constants (1/T<sub>2</sub>) of the |-1/2><-3/2|, |3/2><1/2| coherences (s<sub>1</sub>=410 sec<sup>-1</sup>) and the |1/2><-1/2| coherence (s<sub>2</sub>=100 sec<sup>-1</sup>) were obtained for <sup>23</sup>Na in the albumin solution with the creation time ( $\tau$ ) varied over the range from 0.25 to 20 msec (r<sup>2</sup>=0.997, n=41). The double quantum evolution time ( $\delta$ ) was 0.005 msec, and the relaxation delay (d) was 0.5 sec. The spectral width was 10 kHz, the receiver dead time after the last 90° pulse was 65  $\mu$ sec, and 1024 data points were used. Apparent values of  $\omega \tau_c$  and  $\tau_c$  were calculated to be 1.8 and 5.5×10<sup>-8</sup> sec, respectively. The maximum intensity of the signal was obtained at the creation time of 4-5 msec.

Since  $^{23}$ Na in the glycerol solution is in the condition of extreme narrowing ( $\omega\tau_c$  < <1) at 25°C,  $^{23}$ Na in the glycerol solution showed no significant signal from the double-quantum coherence and showed an artifact of dispersion-like line shape with ca. 0.2% of the intensity obtained by the one-pulse sequence (d-90°-acquire). A single transverse relaxation rate constant (237±5 sec<sup>-1</sup>) obtained for  $^{23}$ Na in the glycerol solution by the conventional spin-echo sequence with the

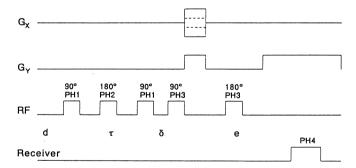
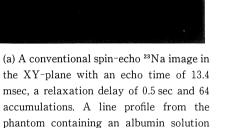


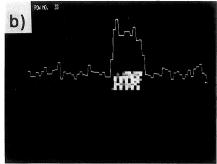
Fig. 1. Pulse sequence for double-quantum filter sodium imaging in the XY-plane without Z -direction slicing. Non-selective  $90^\circ$  and  $180^\circ$  pulses were  $43~\mu{\rm sec}$  and  $86~\mu{\rm sec}$ , respectively. The creation time  $(\tau)$  was 4 msec, the evolution time  $(\delta)$  5  $\mu{\rm sec}$ , the echo time (e) 13.4 msec, and the relaxation

delay 0.2 sec. The phase cycle for the double–quantum filter was as follows: PH1=  $(0^{\circ}, 180^{\circ}, 90^{\circ}, 270^{\circ}, 90^{\circ}, 270^{\circ}, 180^{\circ}, 0^{\circ})_{4}$ , PH2=  $(0^{\circ}, 180^{\circ}, 90^{\circ}, 270^{\circ}, 90^{\circ}, 270^{\circ}, 180^{*}, 0^{\circ})_{2}$   $(180^{\circ}, 0^{\circ}, 270^{\circ}, 90^{\circ}, 270^{\circ}, 90^{\circ}, 0^{\circ}, 180^{\circ})_{2}$ , PH3= $(0^{\circ})_{4}(90^{\circ})_{4}$   $(180^{\circ})_{4}(270^{\circ})_{4}$ , PH4= $(0^{\circ})_{2}(180^{\circ})_{2}(90^{\circ})_{2}(270^{\circ})_{2}$   $(180^{\circ})_{2}(0^{\circ})_{2}(270^{\circ})_{2}(90^{\circ})_{2}$ .



(right side) and a glycerol solution (left

side) is also shown. (b) A double-quantum



filter <sup>23</sup>Na image in the XY-plane with a creation time of 4 msec, an evolution time of 5  $\mu$ sec, an echo time of 13.4 msec and 256 accumulations. The level of suppression of the <sup>23</sup>Na in the glycerol solution is evident in the line profile.

echo time varied over the range from 0.01 to 5 msec (regression coefficient  $\pm$ standard error,  $r^2 = 0.991$ , n=21).

 $^{23}$ Na images in the XY-plane were obtained with a field of view of  $7\times7$  cm (1.5 gauss/cm) across a  $64\times64$  matrix. The radiofrequency and gradient pulse sequences used for double-quantum filter sodium imaging are shown in Fig. 1. To avoid contamination of the signal from  $^{23}$ Na in the extreme narrowing condition, the  $180^{\circ}$  pulse width was measured carefully with an accuracy of  $0.2~\mu$ sec in every experiment.

Figure 2a shows a conventional spin-echo image obtained with an echo time of 13.4 msec. A line profile from the phantom is also shown. <sup>23</sup>Na is visible in both the albumin solution and the qlycerol solution. When we applied the double-quantum filter to the phantom with a creation time of 4 msec, the image from <sup>23</sup>Na in the glycerol solution was suppressed completely and only the signal from <sup>23</sup>Na in the albumin solution was observed in the image (Fig. 2b). One of disadvantage of the double-quantum filter might be low signal-to-noise ratio (1/5-1/10) compared with the conventional spin-echo sequence.

This result indicates the possibility of mapping the correlation time of <sup>23</sup>Na in biological systems. In the perfused rat mandibular gland, <sup>23</sup>Na in the intracellular and interstitial spaces have different relaxation rates. Since the detectable magnetization under the on-resonance condition is in proportion to  $(e^{-s_2}\tau - e^{-s_1}\tau) \cdot e^{-s_{dq}}\delta$ , we may be able to selectively image <sup>23</sup>Na that has a specific correlation time.

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#### References

- A. Bax, R. Freeman, and S.P. Kempsell: Natural abundance <sup>13</sup>C-<sup>13</sup>C coupling observed via double-quantum coherence, J Am Chem Soc, 102: 4849-4851, 1980
- U. Piantini, O.W. Sørenses, and R.R. Ernst: Multiple quantum filters for elucidating NMR coupling networks, J Am Chem Soc, 104: 6800 -6801, 1982
- A.J. Shaka and R. Freeman: Simplification of NMR spectra by filtration through multiplequantum coherence, J Magn Reson, 51:169-173, 1983
- 4) G. Jaccard, S. Wimperis, and G. Bodenhausen: Multiple-quantum NMR spectroscopy of S=3/ 2 spins in isotropic phase: A new probe for multiexponential relaxation, J Chem Phys, 85:6282-6293, 1986
- 5) W.D. Rooney, T.M. Barbara, and C.S. Springer Jr.: Two-dimensional double-quantum NMR

- spectroscopy of isolated spin 3/2 systems: <sup>23</sup>Na examples, J Am Chem Soc 110: 674-681, 1988
- 6) J. Pekar, P.F. Renshaw and J.S. Leigh Jr.: Selective detection of intracellular sodium by coherence-transfer NMR, J Magn Reson, 72: 159-161, 1987
- 7) Y. Seo, M. Murakami, E. Suzuki, et al.: NMR characteristics of intracellular K in the rat salivary gland: A <sup>39</sup>K NMR study using double -quantum filtering, Biochemistry, 29: 599-603, 1990
- 8) T. Hiraishi, Y. Seo, M. Murakami, et al.: Detection of biexponential relaxation in intracellular K in the rat heart by double quantum <sup>39</sup>K NMR, J Magn Reson, in press, 1990
- 9) J. Pekar and J.S. Leigh Jr.: Detection of biexponential relaxation in sodium-23 facilitated by double-quantum filtering, J Magn Reson, 69: 582-584, 1986