

A low-noise stimulated Raman scattering CMOS imager using high-speed lock-in pixels

De Xing Lioe¹, Kamel Mars¹, Sang Man Han¹, Taishi Takasawa¹, Keita Yasutomi¹, Keiichiro Kagawa¹, Mamoru Hashimoto² and Shoji Kawahito¹

¹ Research Institute of Electronics, Shizuoka University
3-5-1 Johoku, Hamamatsu, 432-8011 Japan

² Graduate School of Engineering Science, Osaka University
Toyonaka, 560-8531 Japan

E-mail: lioe@idl.rie.shizuoka.ac.jp

Abstract We demonstrate a low-noise stimulated Raman scattering (SRS) CMOS imager using high-speed lock-in pixels. To detect small SRS signal under an extremely large offset due to probing laser signal of which the ratio is 10^{-5} , a lock-in pixel using a high-speed demodulator, low-pass filter, and multiple-sampling amplifier is designed. The residual offset and low-frequency noise are reduced by two steps; one is sampling phase adjusting in the multiple-sampling amplifier and the other is a double modulation technique. The double modulation effectively reduces the offset and low-frequency noise, or $1/f$ noise. A SRS CMOS imager chip for the proof of concept is implemented and a successful operation is demonstrated. The detectable AC signal level under the large DC offset is measured to be smaller than ratio of 10^{-4} . This finding suggests that the detection of a very small SRS signal from a biological sample is feasible using this technique.

Keywords: stimulated Raman scattering (SRS), lock-in pixels, noise reduction

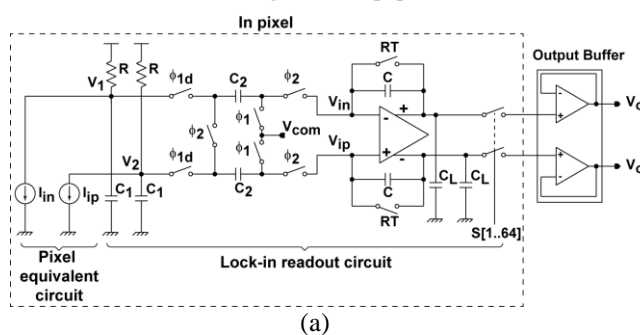
1. Introduction

Recent advances in coherent spectroscopy have made stimulated Raman scattering (SRS) emerged as a label and nonresonant-background-free imaging technique. SRS non-invasively offers imaging contrast mechanism of biological samples [1][2]. SRS operates with two laser beams, pump beam (ω_p) and Stokes beam (ω_s), to interact with the sample, which is in contrast with spontaneous Raman scattering which uses only a single laser beam. Once the frequency difference, $\Omega = \omega_p - \omega_s$, also known as Raman shift, coincides with a particular molecular resonance frequency, stimulated excitation occurs. With the resulting pump signal exists at the same wavelength as the incident laser source, the weak SRS signal which is 10^{-4} to 10^{-5} times smaller than the input signal is hardly distinguishable. Despite this, high-speed modulation lock-in pixels [3][4] can be used to extract the SRS signal. This paper introduces the SRS

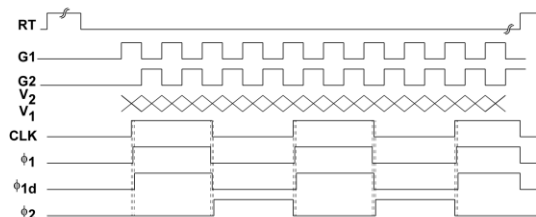
acquisition and noise reduction techniques.

2. Imager and the Operation

The architecture of the lock-in pixel is shown in Fig. 1(a), which consists of pixel equivalent circuit of the modulator, low pass filter, sample and holding circuitry, and multiple-sampling amplifier. Fig. 1(b) shows the timing operation of the lock-in pixel circuit. Initially, the amplifier is reset through the switches RT, in which all other clocks are in OFF state. Switches ϕ_1 are then turned ON to allow capacitors C2 to be set to the reference Vcom level. After a small delay, switches ϕ_{1d} are turned ON in order for capacitors C2 to be charged. Both ϕ_1 and ϕ_{1d} are then turned OFF one after another before switches ϕ_2 are turned ON for charges in C2 to be transferred to capacitors C for integration.

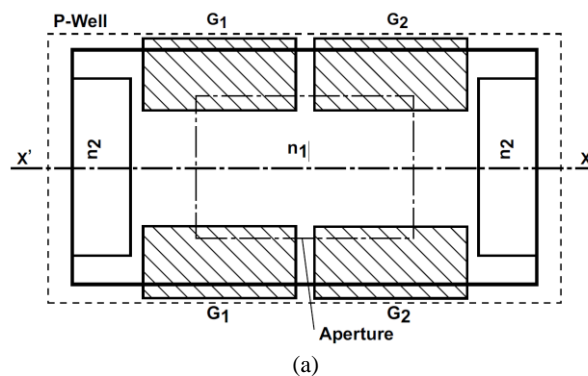


(a)

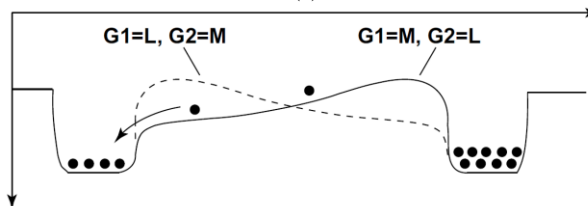


(b)

Fig. 1: Lock-in pixel (a) schematic diagram
(b) timing diagram



(a)



(b)

Fig. 2: Lateral Electric Field Charge Modulator (a) top view
(b) potential profile

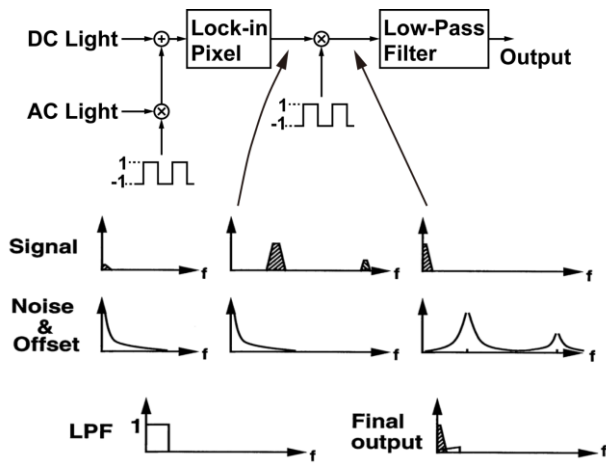


Fig. 3: Double modulation methodology

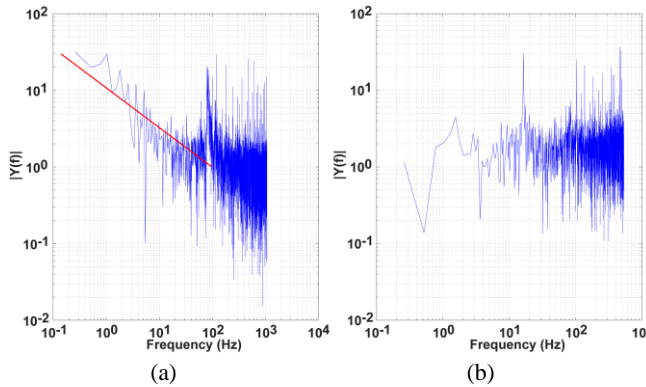


Fig. 4: Effect of noise reduction (a) without double modulation (b) with double modulation

Fig. 2(a) shows the lateral electric field modulator (LEFM), which uses two gates, G_1 and G_2 , for applying lateral electric field to control the horizontal direction of electron flow, rather than photo charge draining. The modulation is performed by applying a medium voltage to one gate and low voltage to another, as depicted in Fig. 2(b). A strong electric field is created in this way which leads to very high transfer rate of lower than 100ps.

3. Noise Reduction Techniques and Results

The reduction of residual offset and low-frequency noise are performed in two steps. The first is the sampling phase adjustment where the phase delay of the demodulated pixel output signal and the readout clock ϕ_1 , ϕ_{1d} and ϕ_2 are carefully adjusted to ensure that the offset signal is cancelled and to achieve high dynamic range. The differential output of the amplifier should be equal to zero during the absence of signal.

A double modulation noise reduction technique is the next step to overcome the low-frequency noise, as shown in Fig. 3. Initially, AC light signal is multiplied by the square wave signal. After this first modulation, the signal is transposed to a higher frequency while the noise and offset components remained. The signal is then amplified and upon second modulation, demodulated back to the baseband. At the same time, the noise component is transposed to high frequency. This technique effectively separate the signal and noise to low and high-frequency components, respectively. The higher frequency noise components are then removed by low-pass filter.

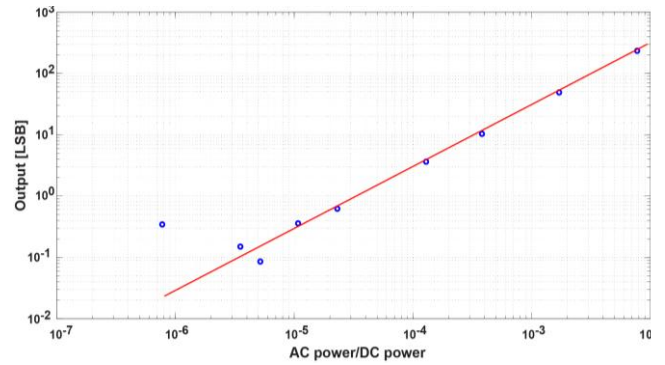


Fig. 5: Ratio of AC power / DC power

4. Experimental Results

A SRS CMOS image sensor chip has been fabricated using $0.11\mu\text{m}$ CMOS image-sensor process technology to evaluate the performance. Fig. 4(a) and (b) illustrate the Fast Fourier Transform (FFT) of the measured signal noise with and without double modulation, respectively. Noise is evident in the low-frequency region without double modulation, while the noise is greatly reduced with double modulation technique applied.

Fig. 5 shows the plot of an average value from 6 measurements of the ratio of the AC (pump) signal to the DC (Stoke) signal. Ratio of approximately 10^{-5} is achieved, which is vital in very weak SRS signal detection.

5. Conclusion

The paper presents the verification of the SRS CMOS imager prototype. The performance of the noise reduction techniques was developed and verified. The imager achieved the AC signal to DC signal power ratio of 10^{-5} , which is an essential region to obtain the SRS signal, and consequently further improves the dynamic range of the SRS CMOS imager.

References

- [1] Christian W. Freudiger, Wei Min, Brian G. Saar, Sijia Lu, Gary R. Holtom, Chengwei He, Jason C. Tsai, Jing X. Kang, and X. Sunney Xie, "Label-Free Biomedical Imaging with High Sensitivity by Stimulated Raman Scattering Microscopy" *Science* 322 (2008), 1857
- [2] P. Nandakumar, A. Kovalev, and A. Volkmer, "Vibrational imaging based on stimulated Raman scattering microscopy" *New Journal of Physics* 11(2009) 033026
- [3] Z. Li, S. Kawahito, K. Yasutomi, K. Kagawa, J. Ukon, M. Hashimoto, Hirohiko Niioka, "A time-resolved CMOS image sensor with draining-only modulation pixels for fluorescence lifetime imaging", *IEEE Trans. Electron Devices*, vol. 59, no. 10, pp. 2715-2722, Oct. 2012.
- [4] S. Kawahito, G. Baek, Z. Li, S.-M. Han, M.-W. Seo, K. Yasutomi, K. Kagawa, "CMOS Lock-in Pixel Image Sensors with Lateral Electric Field Control for Time-Resolved Imaging", *Proc. Int. Image Sensor Workshop*, 2013.