User Guide:

Single-Voxel Spectroscopy semiLASER

Siemens syngo VD and VE platforms

Dinesh Deelchand Sept 2016 Version 1.0

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1. Sequence description

The svs_sLASER sequence presented here is designed for ¹H localized MR spectroscopy in the brain at high and ultra high magnetic fields such as 3T and 7T. sLASER package offers a relatively short T_E sequence that provides full intensity signal similar to PRESS, but optimized to minimize/eliminate unwanted coherences and to generate high quality spectra that enable the reliable quantification of a profile of 10-20 neurochemicals from selected brain regions. Additionally, the sequence is designed to reduce chemical shift displacement errors, which are more severe at higher field strength due to increased chemical shift dispersion.

The current svs_sLASER sequence [1] (Figure 1) is based on the modified semi-adiabatic localization by adiabatic selective refocusing [2], and combines critical features for optimized *in vivo* MRS at high field scanners including:

1- A short T_E (26-30 ms) and slice-selection with pairs of adiabatic 180° pulses in two dimensions for reduced T₂ relaxation effects and reduced J-modulation of coupled spins.

2- High bandwidth slice selective RF pulses for minimal chemical shift displacement.

3- Outer volume saturation (OVS), for more accurate localization and reduced signal contamination from outside the VOI.

4- The VAPOR (**Va**riable **P**ower and **O**ptimized **R**elaxation Delays) technique [3] for excellent water suppression.

5- A loop routine for parameter optimization to maximize spectral quality and minimize signal loss from misadjusted parameters.

6- Single-shot acquisition, i.e. saving of individual FIDs, to enable frequency and phase correction of individual scans and elimination of shots corrupted by subject motion.

A detailed description of the sequence svs_sLASER is found in reference 1. The timing diagram in Figure 1 depicts positioning of the VAPOR and OVS modules with respect to the semi-LASER localization scheme.



Fig. 1. The scheme of the modified semi-LASER sequence. Three pairs of OVS pulses are applied in the X-dimension selected by the slice selective excitation pulse while single pairs of OVS pulses are applied in the Y- and Z-dimensions. [From Öz G & Tkáč, I. Magn Reson Med 2011; 65(4):901-10]

REFERENCES

- Öz G, Tkáč . Short-echo, single-shot, full-intensity ¹H MRS for neurochemical profiling at 4T: Validation in the cerebellum and brainstem, Magn Reson Med 2011; 65:901-10.
- Scheenen TW, Klomp DWJ, Wijnen JP, Heerschap A. Short Echo Time ¹H-MRSI of the Human Brain at 3T With Minimal Chemical Shift Displacement Errors Using Adiabatic Refocusing Pulses, Magn Reson Med. 2008;59:1-6.
- Tkáč I, Starčuk Z, Choi IY, Gruetter R. In vivo ¹H NMR Spectroscopy of Rat Brain at 1 ms Echo Time, Magn Reson Med 1999; **41**: 649-656.

2. Sequence Features

2.1 The Sequence/Special card

Special features of the svs_sLASER are found in the "Sequence/Special" card of the user interface and decribed below.

Routine	Contrast	Resolution	Geometry	System	Ph	iysio	Sequence			
			Comm	ion Spe	cial					
Excitation duration 🔹 🕨 2000 📜 us						Cali	ibration Type	None	•	
		Excitation of	duration:	2000	us		Debug Type	None		
		Relocusing	uulation.	-1000	us					
GOIA/FOC	a 🔹 🕞 GOI/	a/foci 🗌								
				Gradie	nt Ma	ax. Am	plitude 🔹 🕨		40	mT/m

Figure 2: New features included in the Sequence/Special card

The Special features include:

- 1- Settings for the duration of the excitation and refocusing pulses.
- 2- Options for using gradient offset independent adiabatic, e.g. GOIA-HS, GOIA-WURST,
 FOCI and BASSI pulses.
- 3- An option for including an inversion pulse with user selectable inversion time to acquire macromolecule spectra.
- 4- An option for enabling automatic calibration of excitation and VAPOR water suppression flip angle in the MRS VOI (FA AutoCalib). NOTE that this option only works when multiple protocols are setup; refer to the distributed sLASER.exar1 protocols (on VE11B) for more details.
- 5- A calibration loop, which is a very useful tool for optimization of parameters that may degrade spectral quality and cause signal loss is also provided. Those parameters

include the 90° and 180° flip angles, the water suppression (VAPOR) flip angle, VAPOR delays (i.e. 7 and 8) and the inversion time. Selection of the parameter to be optimized is done from the drop-down menu of the 'Calibration Type' (Figure 3). When optimization is completed, the "none" option is selected for acquisition of the final spectral data (Figure 3). Starting values of the 90°, 180° for the calibration procedure are set in the contrast card (Figure 4) while the number of measurements and incremental step size are in the Special Card (Figure 5).

6- A debug loop to acquire VOI and OVS profiles of the selected MRS voxel. The spectra width and number of points are automatically adjusted when this option is enabled.

Routine	Contrast	Resolution	Geometry	System	Physio	Sequence	
			Comm	ion Spec	ial		
Excitatior	duration	> 200	0 📫 us		Cali	bration Type	None 🗸
						Debug Type	None
							Excitation FA
							Refocusing FA
							<vapor fa="" ws=""></vapor>
							<vapor 7="" delay=""></vapor>
							<vapor 8="" delay=""></vapor>
GOIA/FOC	I 🕘 🕨 GOL	a/foci 🗌					<ti></ti>
				Gradien	it Max. Am	plitude 🖪 🕨	40 🗧 mT/m

Figure 3: Calibration options in the Sequence/Special card



Figure 4: Starting values for the FA calibration option in the Contrast/Common card.

2.2 FA calibration

When the "90° Calibration" is selected in the Sequence/Special card, a new field appears for entering incremental steps and number of measurements of the calibration loop. Since the calibration is performed on the water signal, VAPOR water suppression is automatically set to "None" in the Contrast/Common card. This option is only available on VE platform for now.

Routine	Contrast	Resolution	Geometry	System	Physio	Sequence		
			Comm	ion Spec	cial			
Excitation	duration	F 200	0 📫 us		Cali	bration Type	Excitati *	
						Debug Type	None *	
					Increm	ent step size	10 -	deg
					М	easurements	8	
GOIA/FOCI GOIA/FOCI								
				Gradier	nt Max. Am	plitude 🚽 🕒	40 📜	mT/m
			8	3				
Me	asurements	1					20	

Figure 5: Setting of the incremental steps and number of measurements in the Sequence/Special card during 90 Calibration.

Real time monitoring of the calibration loop measurement is done with the "Inline display" feature in the exam card. Examples of the inline display are shown in figures 6 and 8 from in vivo measurements.



Figure 6: Inline display during 90deg calibration of the flip angle in vivo. The minimum and maximum signals are labeled in red with the corresponding flip angle and transmitter reference amplitude. When FA AutoCalib is enabled, the maximum intensity spectrum (denoted here with yellow arrow) will be automatically selected by the system. Yellow dotted line was added for display purpose only in this figure.

When the "VAPOR FA Calibration" is selected in the Sequence/Special card, "VAPOR WS" is automatically enabled. Starting value of the VAPOR FA calibration is entered in the VAPOR flip angle field, and incremental steps in the "Increment step size" field (Figure 7).

Routine	Contrast	Resolution	Geometry	System	Physio	Sequence	
			Commo	n			
		TR 500	0 📜 ms	VAPO	R WS E	nabled	•
	TE 1		8 📜 ms	Water s	uppr. BW	7	0 📜 Hz
	Tota	ITE 2	8 ms				
	Avera	ges	1 ÷	VAPOR F	Flip angle	6	0 📜 deg
Excitation	Flip angle	9	0 🗧 deg				
Excitatio	n Flip angle	0		90			180
Douting	Contract	assilution Co	omotov Custor	Dhusia	Coquer		
Koutine	Contrast r	desolution de	Common S	pecial	Sequent	Le	
Excitation	duration 🖪 🗖	2000	us	Cal	libration Ty	vpe VAPOR	
					Debug Ty	vpe None	
				Increm	nent step s	ize	2 🔆 deg
				M	1easureme	nts	5
goia/foci	d ► GOIA/F	oci					
٥v	'S slab X 🚽 !	120	mm Grad	lient Max. An	nplitude 4	•	40 📜 mT/m

Figure 7: Setting the VAPOR flip angle Calibration and VAPOR bandwidth in the Contrast/Common card and incremental steps and number of measurements in the Sequence/Special card.



Figure 8: Inline display during calibration of the VAPOR flip angle in vivo. The flip angles producing the smallest and largest water signals are labeled in red. When FA AutoCalib is enabled, the minimum water signal spectrum (denoted here with yellow arrow) will be automatically selected by the system.

2.3 Saving single-shot FIDs

The capability of saving individual averages and the summed spectrum (Figure 9) is also available. With the VAPOR option selected,"Save single averages" saves both the individual averages as well as the summed spectrum.

Routine	Contrast	Resolution	Geometry	System	Physio	Sequence	
Coi	ils Miscel	laneous Adj	ustments	Adjust Volum	ne pTx	Volumes	Tx/Rx
Pos	itioning mor	le RFF			Save	uncombine	d
103					5470	uncombine	
Table posi	tion H	• 0	mm				
		0 0 T					
	MSMA	S - C - I	·				
	Sagittal	R >> L	T		Save sir	ngle average	s 🗸
	Coronal	A >> P	•	AutoAlig	n		·
т	ransversal	F >> H		Coil Select	Mode	Default	*

Figure 9: Save single averages option is located in the System/Miscellaneous card.

2.4 Gradient-modulated RF pulses

The default refocusing pulses in sLASER are an HS4-R25 adiabatic full passage pulse which requires a B₁ of ~25 μ T (when GOIA/FOCI option in the Sequence/Special card is disabled). If the B₁ is limited or if one wants to further reduce the echo-time of sLASER, gradient-modulated RF pulses can be used. Four types of refocusing RF pulse are available (Figure 10) and these are GOIA-HS, GOIA-WURST, FOCI and BASSI RF pulses.

For GOIA RF pulses, the bandwidth at 1ms, modulation of the HS pulse and gradient factor can be entered in the Sequence/Special card (Figure 11). For FOCI, only the bandwidth can be changed. The bandwidth of BASSI pulse (and other RF parameters) is hard-coded to be 45 kHz at 1 ms duration. This pulse was optimized for 4.5 ms duration at a required B₁ of 15 μ T.

<u>NOTE</u>: Before using these type of refocussing RF pulses, please run Bloch simulator based on the available B_1 field of your RF coil to determine the required duration/bandwidth/gradient factor.

Routine Contrast	Resolution Geometry	y System Physio Sequence
	Con	nmon Special
Excitation duration	> 2000 ÷ us	Calibration Type None 🔻
АҒР Туре	e GOIA 🗸	Debug Type None 🔻
Bandwidth_1ms	GOIA-HS GOIA-WURST FOCI BASSI	
goia/foci 🚽 🕨 goia/	Foci 🗹	
		Gradient Max. Amplitude и 🕨 40 🗧 mT/m

Figure 10: Gradient-modulated refocusing RF pulse options in the Sequence/Special card

Routine	Contrast	Resolution	Geometry	System	Physio	Sequence	
			Comm	ion Spe	cial		
Excitatior	duration	P 200	0 📜 us		Cali	ibration Type	None 🔻
	AFP Ty	vpe GOIA	•			Debug Type	None 🔻
Bandw	vidth_1ms 4	► 5	0 📜 kHz				
Bandwidth_1ms: 50 HSn modulation: 16 Gradient factor: 90				kHz %			
GOIA/FOC	I 🔹 🕨 GOI/	a/foci 🗹					
				Gradier	nt Max. Am	plitude 🖪 🕨	40 📮 mT/m

Figure 11: Bandwidth, HS modulation or gradient factor to define the gradient-modulated RF pulse in the Sequence/Special card

2.5 Water references scans

A new feature available on sLASER is the possibility to save the water reference scans and metabolites data together as individual FIDs within one protocol. The water scans includes the water reference scans for eddy-current correction (set VAPOR WS to "Only RF off" in Contrast/Common card) and for quantification (set VAPOR WS to "None" in Contrast/Common card).

This "Water Ref. Scan" option is only visible when VAPOR is enabled (Figure 12) and the "No. of ref. scans" field defines the number of scans to acquire before and after the metabolite spectrum is acquired, as shown in the data structure (Figure 13).

Hz
ms

Figure 12: Bandwidth, HS modulation or gradient factor to define the gradient-modulated RF pulse in the Sequence/Special card



Figure 13: The structure of the water suppressed protocol when Water Ref. Scan is enabled.

2.6 Automatic RF calibration

Another new feature implemented in sLASER is the option to automatically perform the excitation and VAPOR flip angle calibrations. This option only works if four separate protocols are setup as described below (Figure 14). Note that on VE11B, sLASER.exar1 file are provided for these protocols:

1) sLASER_LW

<u>Contrast/Common card</u>: Excitation Flip angle = 90 deg, Measurements = 1, Averages = 1, VAPOR WS = None; FA AutoCalib = off <u>Sequence/Common card</u>: Preparation scans = 1, Delta frequency = 0 ppm <u>Sequence/Special card</u>: Calibration type = None

2) sLASER_90Calib

<u>Sequence /Special card</u>: Calibration type = Excitation FA, Increment step size = 10 deg, Measurements = 8; FA AutoCalib = On

<u>Contrast/Common card</u>: Excitation Flip angle = 50 deg, Averages = 1, VAPOR WS = None.

<u>Sequence/Common card</u>: Preparation scans = 1, Delta frequency = 0 ppm

3) sLASER_WSCalib

<u>Sequence/Special card</u>: Calibration type = VAPOR WS FA (this automatically sets VAPOR WS = enabled), increment = 2 deg, Measurements = 8, FA AutoCalib = on <u>Sequence/Common card</u>: Preparation scans = 1, Delta frequency = -2 ppm <u>Contrast/Common card</u>: Excitation Flip angle = 90 deg, Averages = 1, VAPOR WS = Enabled, VAPOR flip angle = 52

4) sLASER_WS64

Sequence/Special card: Calibration type = None, FA AutoCalib = On

<u>Sequence/Common card</u>: Preparation scans = 1, Delta frequency = -2 ppm <u>Contrast/Common card</u>: Excitation Flip angle = 90 deg, Measurements = 1, Averages = 64 (or 32)

<u>IMPORTANT</u>: Set up steps #2-4 such that the "Measurement Parameters" is copied from step 1 (Figure 14).

New Tree » Dinesh » MRS » sL	ASER
sLASER_LW	00:15
sLASER_90Calib	00:50
sLASER_WSCalib	01:10
sLASER_WS64	06:10

Figure 14: Protocols required for successfully running the automatic RF calibration

3. Measurement Protocols

This svs_sLASER sequence acquires single shots that can be exported and corrected for frequency and phase shifts prior to summing. The following common parameters are set as default in the protocol.

- In Contrast card
 - TR = 5000 ms
 - \circ TE1 = 8 ms, TE2 = 11 ms and TE3 = 9 ms, i.e. total TE = 28 ms
 - Water s. BW = 70 Hz
- In Sequence/Special card:
 - Excitation duration = 2000 μs
 - Refocussing duration = 4000 μs
 - FOCI/GOIA = Off
- In Sequence/Common card:
 - Bandwidth = 6000 Hz

Two B_0 shimming technique can be used with sLASER: FAST(EST)MAP or system 3D-GRE shimming routines as described below.

3.1 Using FAST(EST)MAP shim

The following workflow includes use of the FASTESTMAP shimming procedure based on the original work by Rolf Gruetter and Ivan Tkáč (Gruetter and Tkáč, Magn. Reson. Med. 43:319-324, 2000). The FASTESTMAP sequence is available from Edward J. Auerbach, Center for Magnetic Resonance Research (CMRR), Minneapolis, MN, USA. Create the protocols, as described below, and use them as suggested in the workflow (next section).

• Fastestmap-Linear_3

- <u>Contrast/Common card</u>: Tau = 5 ms
- <u>System/Adjustments card</u>: B0 Shim mode = Tune Up
- <u>Sequence/Special card</u>: Type of fit = Linear 3-proj, Bar FoV = 300 mm, Bar thickness = 5 mm, multi-echo acquisition = On, Number of echoes = 8, Inversion pulse = Off
- Fastestmap-Full_6
 - <u>Contrast/Common card</u>: Tau = 5 ms

- <u>System/Adjustments card</u>: B0 Shim mode = Advanced
- <u>Sequence/Special card</u>: Type of fit = Full 6-proj, Bar FoV = 300 mm, Bar thickness = 5 mm, multi-echo acquisition = On, Number of echoes = 8, Inversion pulse = Off
- Fastestmap-Linear_6
 - <u>Contrast/Common card</u>: Tau = 10 ms
 - <u>System/Adjustments card</u>: B0 Shim mode = Advanced
 - <u>Sequence/Special card</u>: Type of fit = Linear 6-proj, Bar FoV = 300 mm, Bar thickness = 5 mm, multi-echo acquisition = Off, Inversion pulse = Off

3.2 Using system 3D-GRE shim

To use the system 3D-GRE B_0 shim option, the "B0 Shim mode" in the System/Adjustments card must be set to "Brain" (Figure 15) in the sLASER LW protocol (seciton 2.6). After positioning the VOI in the LW protocol, the system will automatically adjust the first- and second-order shims in one or two iterations before acquiring the sLASER data.

Routine Contrast Res	olution Geometry	System Physio	Sequence	
Coils Miscellan	eous Adjustments	Adjust Volume	pTx Volumes	Tx/Rx
B0 Shim mode Brain	•	B1 Shim mode	TrueForm	
Adjustment Tolerance Aut	to 🔻			
Adjust with	body coil 🗌			
Confirm freq. ad	djustment			
		As	ssume Dominar	nt Fat 📃
			Assume Sili	icone
		in and to Design it		

Figure 15: Set system B0 shim mode to Brain with sLASER

4. Measurement Workflow

The suggested MRS measurement workflow when using either FAST(TEST)MAP or 3D-GRE shim are described below.

- 1. Run Localizer (3 Plane Scout)
- 2. Run MPRAGE
 - a. Extract/Recon images into two other orientations, i.e. transverse and sagittal
 - b. Load all 3 images in the Exam card
 - c. Start "Inline display" and keep it open!

If FAST(EST)MAP shim is used:

- 3. Open Fastestmap-Linear_3
 - a. Position voxel in region-of-interest
 - **b.** Click "Apply".
- 4. Fastestmap-Full_6
 - a. Copy Measurement Parameters from previous step.
 - b. Wait until the scan has ended.
 - i. Press "Ctrl + Esc" buttons
 - ii. Select "Set FASTMAP Shims"
 - iii. Type "a", then Press "Enter"
 - 1. Wait for the blue screen to automatically close
 - c. Click "Apply"
- 5. Fastestmap-Full_6 (repeating the second order shimming step improves linewidth)
 - a. Copy Measurement Parameters from previous step.
 - b. Wait until the scan has ended.
 - i. Press "Ctrl + Esc" buttons
 - ii. Select "Set FASTMAP Shims"
 - iii. Type "a", then Press "Enter"
 - 1. Wait for the blue screen to automatically close
 - c. Click "Apply"

6. Fastestmap-Linear_6

- a. Copy Measurement Parameters from previous step.
- b. Wait until the scan has ended.
 - i. Press "Ctrl + Esc" buttons

- ii. Select "Set FASTMAP Shims"
- iii. Type "a", then Press "Enter"
 - 1. Wait for the blue screen to automatically close
- c. Click "Apply"

7. sLASER_LW

- a. Copy Measurement parameters from Fastestmap-Linear_6.
- b. Wait until the scan has ended.
 - i. Press "Ctrl + Esc" buttons
 - ii. Select "Set FASTMAP Shims"
 - iii. Type "a", then Press "Enter"
 - 1. Wait for the blue screen to automatically close
- c. Click "Apply"

Is the gray vertical line centered on the water peak (green line)? If not, adjust the frequency in Options/Adjustment/Frequency Tab and rerun **sLASER_LW**

d. Goto Step #9 below

If 3D-GRE shim is used:

8. sLASER_LW

a. Position voxel in region-of-interest in 3D MPRAGE images

 $3D B_0$ maps will be acquired in one or two iterations and the shims will be automatically set

b. Click "Apply".

Is the gray vertical line centered on the water peak (green line)? If not, adjust the frequency in Options/Adjustment/Frequency Tab and rerun sLASER_LW

- c. Goto Step #9 below
- 9. These protocols should start automatically: sLASER_90Calib, sLASER_WSCalib and sLASER_WS64
- 10. Run Localizer again (to ensure subject has not moved)
- 11. Done!

5. Water linewidth estimation

The linewidth of the water peak after shimming can be estimated as follows:

- a. Start Spectroscopy Task Tab by selecting Applications / Spectroscopy from top of the screen
- b. Go to Spectroscopy Tab
- c. Load filename containing "sLASER_LW" from the Patient Browser
- d. Go to Protocol Open, then Select "SLASER_waterLW"
- e. The value next to the W (in yellow) will be the estimated linewidth in Hz (Figure 16).



Figure 16: Estimating water linewidth on the scanner

6. Installation

Installation of this package should be performed by an experienced user. For installation you need to access the file-system of the scanner ("Advanced User" functionality), which requires a password. Please contact the local application support for this information.

Please follow the installation steps below:

- Login as "Advanced User".

- Insert your medium in the appropriate drive on the host computer.

- Open "Windows explorer" and find the file "install_slaser.bat" on the inserted medium.

- Double click on the file name to start the installation (note: there is no need to copy the file to the system).

- The window below displays progress of the installation.

- Monitor the installation and follow the displayed messages.

- When the message "Done!" is displayed, close the window by pressing any key on the keyboard (Figure 17).

The installation automatically copies the following files to the appropriate folders on the system:

0	Sequence:		
	svs_slaser_dkd.dll	to	C:\medcom\MriCustomer\Seq\
	libsvs_slaser_dkd.so	to	C:\medcom\MriCustomer\Seq\
0	Ice programs:		
	dkdIceSpecSEAD2.dll	to	C:\medcom\bin\
	dkdIceSpecSEAD2.evp	to	C:\medcom\bin\
	dkdIcePrgSpectro2.ipr	to	C:\medcom\bin\
	libdkdlceSpecSEAD2.so	to	C:\medcom\MCIR\Med\lib\
0	RF pulse:		
	extrf_sead.dat	to	C:\medcom\MriCustomer\Seq\
0	Processing file:		
-	\$\$SI ASER water! W pro	to	C:\medcom\MriCustomer\MrSpec\E\/2Protocols\
	φφοιλοεκ_wateriev.pro	10	

Administrator: C:\Windows\system32\cmd.exe - install_slaser.bat
G:\sLASER_Oct2016>install_slaser.bat
Installer for sLASER on VE platform Dinesh Deelchand, Oct 2016
*** Installing sequence files: svs_slaser_dkd.dll success! libsvs_slaser_dkd.so success! extrf_sead.dat success!
Some system components and the MARS must be restarted in order to properly install the ICE binaries. Before continuing, please make sure no scans are running and note that the system will be unusable for at least 10 minutes while the MARS reboots.
If you continue, the MARS will be rebooted NOW.
Are you sure you want to continue [y/n]? Are you sure you want to continue [y/n]? y
Rebooting MARS ssh root@mars reboot rem attempt to unload the PAS and restart Cocos to hopefully unlock dlls Waiting 10 seconds for MARS shutdown
**** Installing ICE program files: dkdIcePrgSpectro2.ipr success! dkdIceSpecSEAD2.evp success! dkdIceSpecSEAD2.dll success! libdkdIceSpecSEAD2.so success!
*** Installing linewidth protocol \$\$SLASER_waterLW.pro success!
Please import the sLASER protocols (i.e. sLASER.exar1 located in the protocol fo lder) using the Dot Cockpit.
Done!
Press any key to continue

Figure 17: Installation screen when using install_slaser.bat file