Complexity of cardiac signals for predicting changes in alpha-waves after stress in patients undergoing cardiac catheterization

Supplementary Information

Hung-Chih Chiu^{1,*}, Yen-Hung Lin^{2,*}, Men-Tzung Lo^{3,4}, Sung-Chun Tang⁶, Tzung-Dau Wang², Hung-Chun Lu², Yi-Lwun Ho², Hsi-Pin Ma¹ & Chung-Kang Peng^{4,5}

¹Department of Electrical Engineering, National Tsing Hua University, Hsinchu, Taiwan.

²Department of Internal Medicine, National Taiwan University Hospital and National Taiwan University College of Medicine, Taipei, Taiwan.

³Institute of Translational and Interdisciplinary Medicine and Department of Biomedical Sciences and Engineering, National Central University, Chungli, Taiwan

⁴Research Center for Adaptive Data Analysis, National Central University, Taoyuan, Taiwan.

⁵Division of Interdisciplinary Medicine and Biotechnology, Beth Israel Deaconess Medical Center/Harvard Medical School, Boston, Massachusetts, USA.

⁶Department of Neurology, National Taiwan University Hospital and National Taiwan University College of Medicine, Taipei, Taiwan.

Supplement	ary Table S1 Classificat	ion of patients or	n the basis of the pretest variables and the
difference b	etween pre- and post-test	variables.	
			meanNN, Log-meanNN,
			sdNN, Log-sdNN,
	Linear Variable _	Pre-Test	pNN20, Log-pNN20,
			pNN50, Log-pNN50,
			rMMSD, Log-rMMSD,
			LF, Log-LF, HF, Log-HF
			LF/HF, Log-LF/HF
		Difference	meanNN, Log-meanNN,
			sdNN, Log-sdNN,
			pNN20, Log-pNN20,
566			pNN50, Log-pNN50,
ECG			rMMSD, Log-rMMSD,
			LF, Log-LF, HF, Log-HF
			LF/HF, Log-LF/HF
	Non-linear Variable		Slope1-5, Slope6-20,
			Area1-5, Log_Area1-5,
		Pre-Test	Area6-20, Log_Area6-20
			α_1 , Log_ α_1 , α_2 , Log_ α_2
			Slope1-5, Slope6-20,
		Difference	Area1-5, Log_Area1-5,
			Area6-20, Log_Area6-20
			α_1 , Log- α_1 , α_2 , Log- α_2
	Linear Variable	Pre-Test	Alpha activity
EEG		Difference	Alpha activity

Clinical Data

Age, Gender, BMI, MBP(Pre-Test), MBP(Difference), Treatment, Smoking, Diabetes

Mellitus, Hyperlipidemia, Hypertension, EGFR

Neurotransmitter

Serotonin, Orphanin_FQ, Dopamine

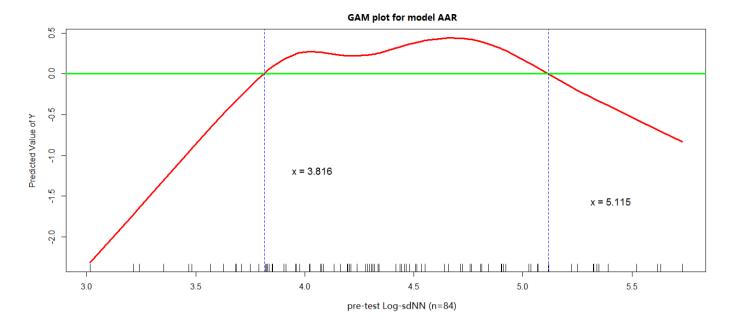
Supplementary	y Table S2 Classification o	of patients accordi	ng to Spearman's rank correlation.			
			meanNN,			
		Due Tret	sdNN, Log-sdNN,			
		Pre-Test	pNN20, Log-pNN20,			
	Linear Variable		LF/HF, Log-LF/HF			
			meanNN,			
		5.10	pNN20, Log_pNN20,			
		Difference	rMMSD,			
ECG			LF/HF, Log_LF/HF			
			Slope1-5,			
		Pre-Test	Area1-5, Log-Area1-5,			
	Non-linear Variable		α_1, α_2			
			Slope1-5,			
		Difference	Area1-5, Log-Area1-5,			
			α ₁ , α ₂			
EEG	Linear Variable	Pre-Test	Alpha activity			
EEG		Difference	Alpha activity			
	Age, Gender, BMI, MBP(Pre	-Test), MBP(Differe	nce),			
Clinical Data	Treatment, Smoking, Diabetes Mellitus, Hyperlipidemia, Hypertension, estimated					
	glomerular filtration rate					
	Neurotransmitter		Serotonin, Orphanin_FQ, Dopamine			

Supplemen	tary Table S3 Clas	sification of meth	ods by using GAMs.				
		Pre-Test	$3.816 \leq \text{Log}_{\text{sdNN}} \leq 5.115$,				
			$0.017 \le \text{pNN50} \le 0.176$,				
			$56.598 \le rMSSD \le 194.443$,				
			$3.81 \leq \text{Log}_r\text{MSSD} \leq 5.348,$				
			$1071.873 \le \text{HF} \le 13485.712$,				
	Linear		$1.229 \le LF/HF \le 1.98$,				
	Variable	Difference	$Log_sdNN \le -0.415$,				
			$-0.164 \le pNN20 \le 0.007$,				
			$-0.142 \le pNN50 \le 0.007$,				
			$-105.641 \le rMSSD \le 4.677,$				
ECG			$LF \le 156.425$ and $LF \ge 13413.356$,				
			${\rm HF} \le 376.933$ and ${\rm HF} \ge 25327.388$				
		Pre-Test	$0.01 \leq \text{Slope6} - 20$,				
			$Log_Area1 - 5 \le 1.007,$				
			$2.183 \le Log_Area6 - 20 \le 2.895$,				
	Non-linear		$0.425 \le \alpha_1 \le 0.756$,				
	Variable		$\alpha_2 \leq 0.737 \text{ and } \alpha_2 \geq 0.876$				
	_	Difference	Area $1 - 5 \le -1.247$ and Area $1 - 5 \ge 0.809$,				
			$Log_Area6 - 20 \le -0.042$ and $Log_Area6 - 2$				
			≥ 0.5				

Supplementary Table S4 | Experimental results of the HRV analysis before *treatment, obtained using Spearman's* rank correlation.

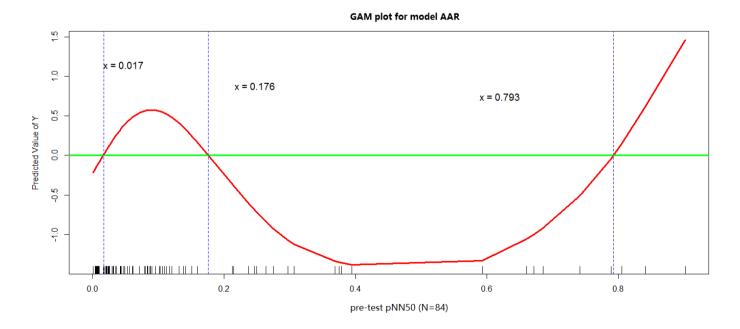
	sdNN	Log-sdNN	pNN50	Log-pNN50	rMMSD	Log-rMMSD	LF	Log-LF	LF/HF	Log-LF/HF
pNN50	0.9157	0.9157								
Log- pNN50	0.9157	0.9157								
rMMSD	0.9129	0.9129	0.8638	0.8638						
Log-rMMSD	0.9129	0.9129	0.8638	0.8638						
LF	0.8364	0.8364			0.845	0.845				
Log-LF	0.8364	0.8364			0.845	0.845				
HF	0.8151	0.8151			0.9092	0.9092	0.9487	0.9487		
Log-HF	0.8151	0.8151			0.9092	0.9092	0.9487	0.9487		
\propto_1									0.8331	0.8331
Log∝ ₁									0.8331	0.8331

Supplementary Table S5 | Experimental results regarding the difference between the pre- and posttest variables obtained using Spearman's rank correlation Log-sdNN LF sdNN Log-LF LF/HF 0.8979 0.8564 rMMSD 0.8320 0.8760 Log- rMMSD HF 0.9486 0.9405 Log-HF 0.9782 Log-LF/HF



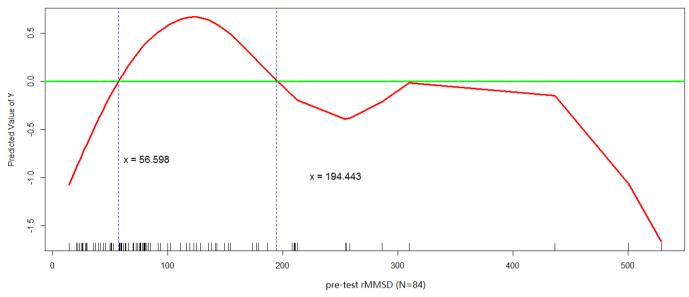
Supplementary Figure S1: GAM plot for modeling the relationship between the change in the alpha wave and pretest log-sdNN. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest log-sdNN and number of patients. GAM techniques can be used to show that

 $pre-test \ Log-sdNN = \begin{cases} 1, & 3.816 \le Log-sdNN \le 5.115 \\ 0, & others \end{cases}$



Supplementary Figure S2: GAM plot for modeling the relationship between the change in the alpha wave and pretest pNN50. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest pNN50 and number of patients. GAM techniques can be used to show that

pre-test pNN50 =
$$\begin{cases} 1, & 0.017 \le \text{pNN50} \le 0.176 \\ 0, & \text{others} \end{cases}$$

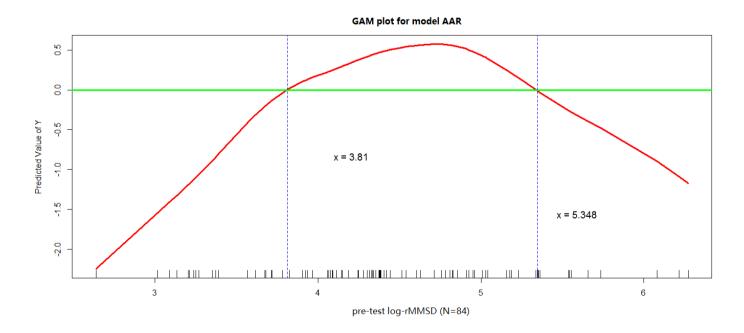


Supplementary Figure S3: GAM plot for modeling the relationship between the change in the alpha wave and pretest

rMMSD. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest rMMSD and number

of patients. GAM techniques can be used to show that

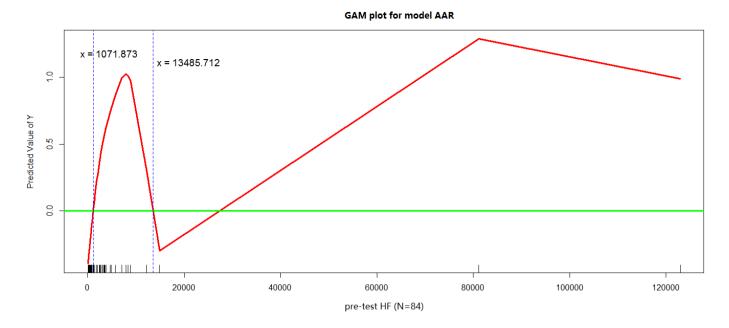
pre-test rMMSD = $\begin{cases} 1, & 56.598 \le \text{rMMSD} \le 194.443 \\ 0, & \text{others} \end{cases}$



Supplementary Figure S4: GAM plot for modeling the relationship between the change in the alpha wave and pretest

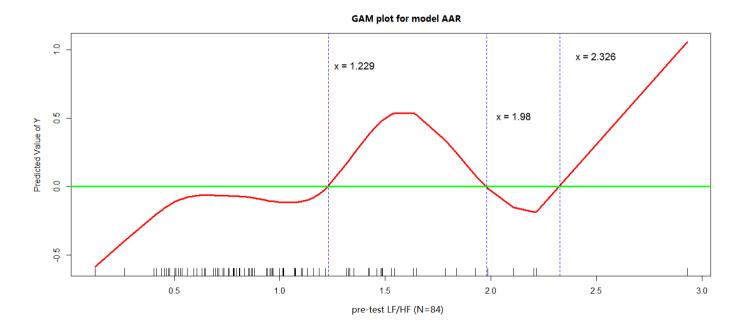
log-rMMSD. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest log-rMMSD and number of patients. GAM techniques can be used to show that

pre-test log-rMMSD =
$$\begin{cases} 1, & 3.81 \le \text{log-rMMSD} \le 5.348 \\ 0, & \text{others} \end{cases}$$



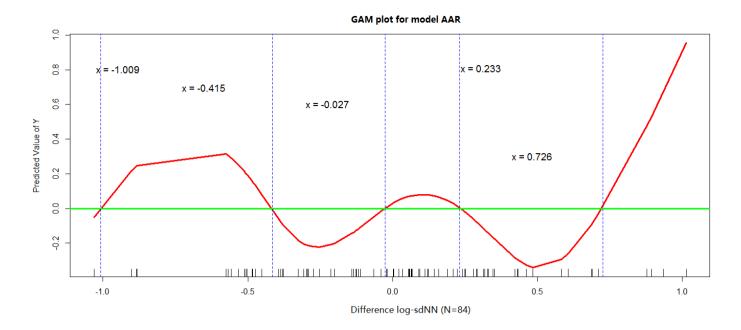
Supplementary Figure S5: GAM plot for modeling the relationship between the change in the alpha wave and pretest HF. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest HF and number of patients. GAM techniques can be used to show that

pre-test
$$HF = \begin{cases} 1, & 1071.873 \le HF \le 13485.712 \\ 0, & others \end{cases}$$



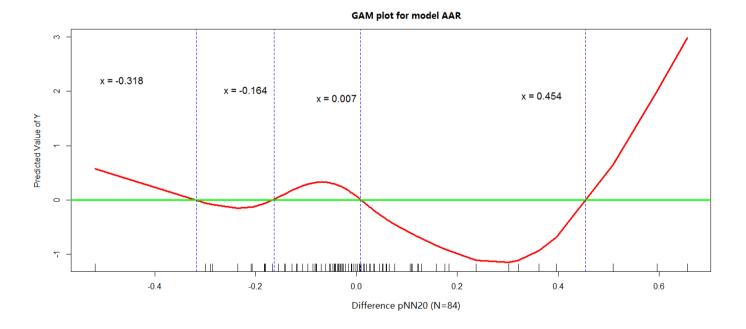
Supplementary Figure S6: GAM plot for modeling the relationship between the change in the alpha wave and pretest LF/HF. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest LF/HF and number of patients. GAM techniques can be used to show that

pre-test LF/HF =
$$\begin{cases} 1, & 1.229 \le \text{LF/HF} \le 1.98\\ 0, & \text{others} \end{cases}$$



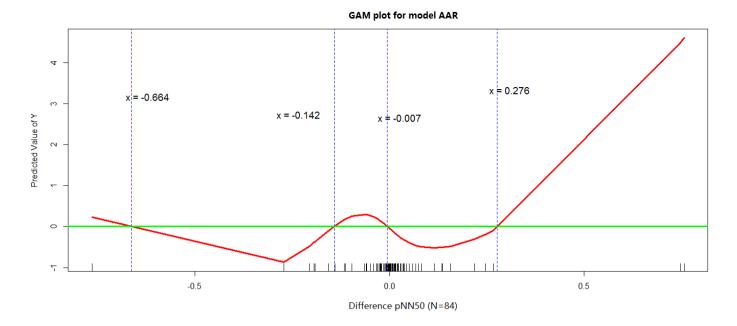
Supplementary Figure S7: GAM plot for modeling the relationship between the change in the alpha wave and Difference log-sdNN. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference log-sdNN and number of patients. GAM techniques can be used to show that

Difference log-sdNN =
$$\begin{cases} 1, & \text{log-sdNN} \leq -0.415 \\ 0, & \text{others} \end{cases}$$



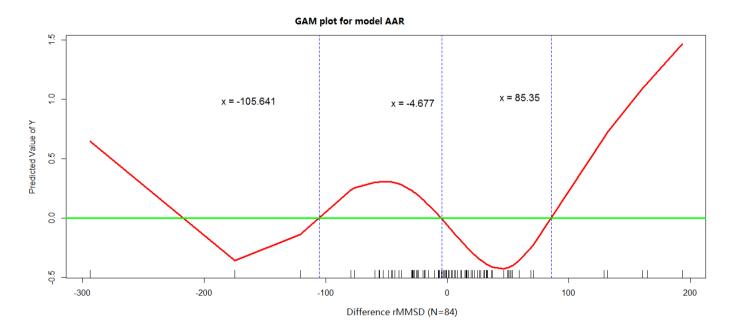
Supplementary Figure S8: GAM plot for modeling the relationship between the change in the alpha wave and Difference pNN20. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference pNN20 and number of patients. GAM techniques can be used to show that

Difference pNN20 =
$$\begin{cases} 1, & -0.164 \le \text{pNN20} \le 0.007 \\ 0, & \text{others} \end{cases}$$



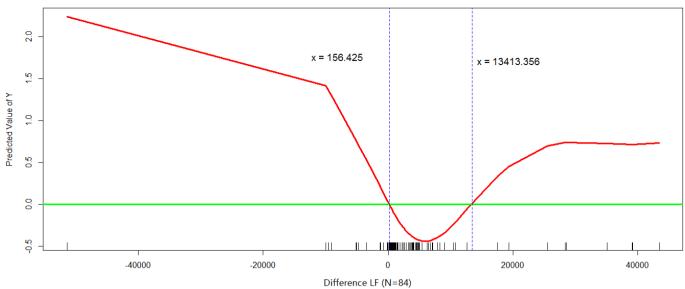
Supplementary Figure S9: GAM plot for modeling the relationship between the change in the alpha wave and Difference pNN50. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference pNN50 and number of patients. GAM techniques can be used to show that

Difference pNN50 =
$$\begin{cases} 1, & -0.142 \le \text{pNN50} \le 0.007 \\ 0, & \text{others} \end{cases}$$



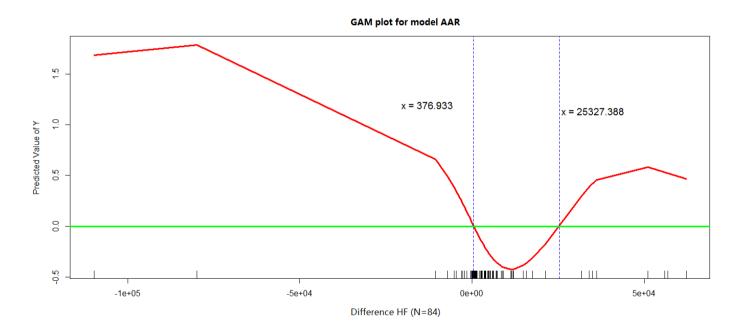
Supplementary Figure S10: GAM plot for modeling the relationship between the change in the alpha wave and Difference rMMSD. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference rMMSD and number of patients. GAM techniques can be used to show that

Difference rMMSD =
$$\begin{cases} 1, & -105.641 \le \text{rMMSD} \le 4.677 \\ 0, & \text{others} \end{cases}$$



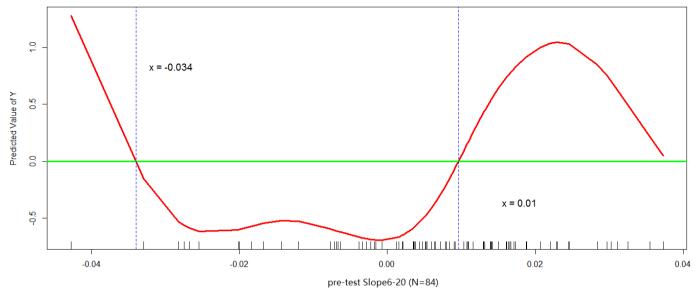
Supplementary Figure S11: GAM plot for modeling the relationship between the change in the alpha wave and Difference LF. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference LF and number of patients. GAM techniques can be used to show that

Difference LF =
$$\begin{cases} 1, & 156.425 \le LF \le 13413.356 \\ 0, & \text{others} \end{cases}$$



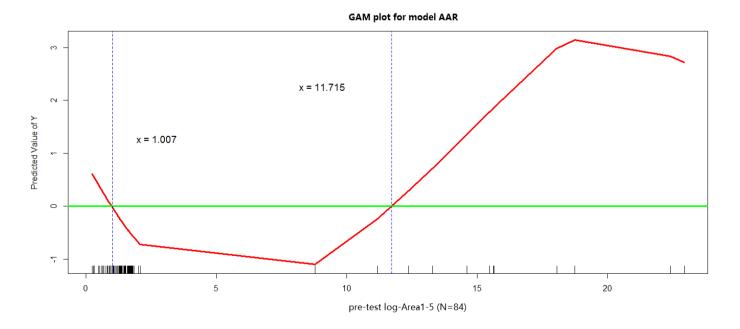
Supplementary Figure S12: GAM plot for modeling the relationship between the change in the alpha wave and Difference HF. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference HF and number of patients. GAM techniques can be used to show that

Difference
$$HF = \begin{cases} 1, & 376.933 \le HF \le 25327.388 \\ 0, & others \end{cases}$$



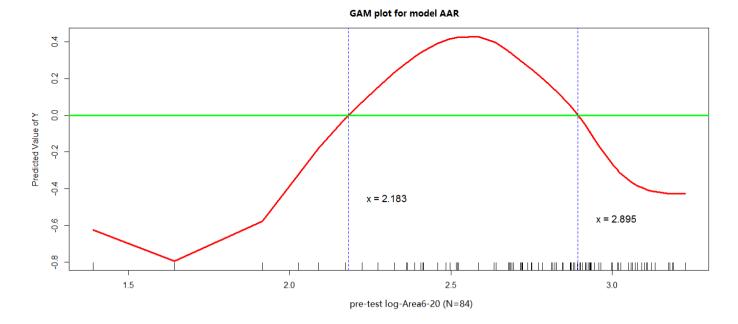
Supplementary Figure S13: GAM plot for modeling the relationship between the change in the alpha wave and pre-test Slope6-20. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest Slope6-20and number of patients. GAM techniques can be used to show that

pre-test Slope6-20 =
$$\begin{cases} 1, & 0.01 \le \text{Slope6-20} \\ 0, & \text{others} \end{cases}$$



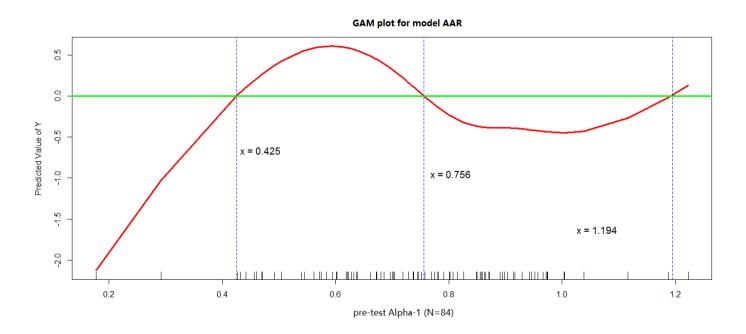
Supplementary Figure S14: GAM plot for modeling the relationship between the change in the alpha wave and pre-test log-Area1-5. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest log-Area1-5 and number of patients. GAM techniques can be used to show that

pre-test log-Area1-5 =
$$\begin{cases} 1, & \text{log-Area1-5} \le 1.007\\ 0, & \text{others} \end{cases}$$



Supplementary Figure S15: GAM plot for modeling the relationship between the change in the alpha wave and pre-test log-Area6-20. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest log-Area6-20 and number of patients. GAM techniques can be used to show that

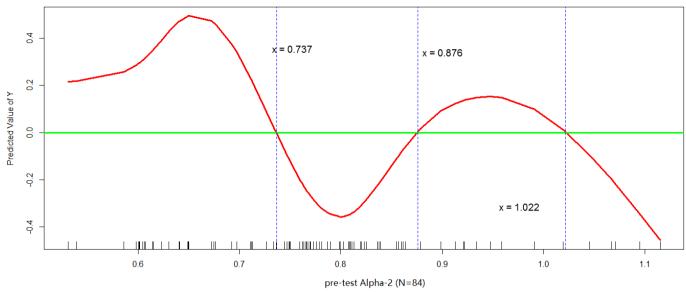
pre-test log-Area6-20 = $\begin{cases} 1, & 2.183 \le \text{log-Area6-20} \le 2.895 \\ 0, & \text{others} \end{cases}$



Supplementary Figure S16: GAM plot for modeling the relationship between the change in the alpha wave and pre-test Alpha-1. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest Alpha-1 and number of patients. GAM techniques can be used to show that

pre-test Alpha-1 =
$$\begin{cases} 1, & 0.425 \le \text{Alpha-1} \le 0.756 \\ 0, & \text{others} \end{cases}$$



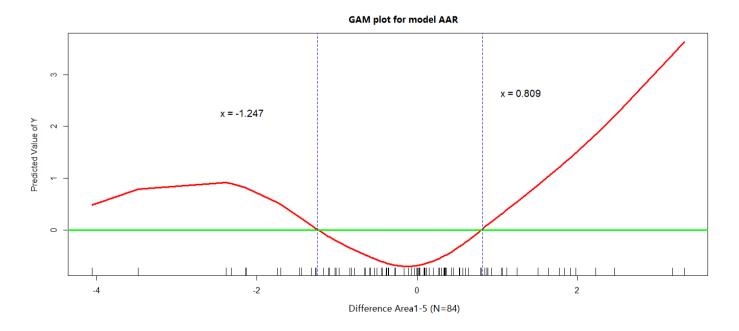


Supplementary Figure S17: GAM plot for modeling the relationship between the change in the alpha wave and pre-test

Alpha-2. The vertical axis represents the difference in alpha activity. The horizontal axis represents the pretest Alpha-2 and number

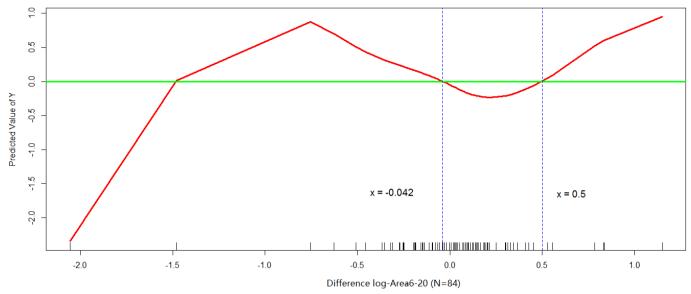
of patients. GAM techniques can be used to show that

pre-test Alpha-2 = $\begin{cases} 1, & \text{Alpha-2} \le 0.737 \text{ and Alpha-2} \ge 0.876 \\ 0, & \text{others} \end{cases}$



Supplementary Figure S18: GAM plot for modeling the relationship between the change in the alpha wave and Difference Area1-5. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference Area1-5 and number of patients. GAM techniques can be used to show that

Difference Area1-5 =
$$\begin{cases} 1, & \text{Area1-5} \le -1.247 \text{ and Area1-5} \ge 0.809 \\ 0, & \text{others} \end{cases}$$



Supplementary Figure S19: GAM plot for modeling the relationship between the change in the alpha wave and Difference log-Area6-20. The vertical axis represents the difference in alpha activity. The horizontal axis represents the Difference log-Area6-20 and number of patients. GAM techniques can be used to show that

Difference log-Area6-20 = $\begin{cases} 1, & \text{Area6-20} \le -0.042 \text{ and } \text{Area6-20} \ge 0.5 \\ 0, & \text{others} \end{cases}$