Supplementary Material

DEMIST: A deep-learning-based task-specific denoising approach for myocardial perfusion SPECT

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S-I. NETWORK ARCHITECTURE

TABLE S-1

NETWORK ARCHITECTURE (CONV. = CONVOLUTIONAL, RELU = RECTIFIED LINEAR UNIT, MAXPOOL = MAX POOLING). BATCH NORMALIZATION WAS USED AFTER EACH OF THE CONVOLUTIONAL AND TRANSPOSED CONVOLUTIONAL LAYER.

Layer	Layer type	Number of filters	Filter size	Stride/Pool size	Input size	Output size
1	Conv.	16	3×3×3	1×1×1	48×48×48×1	48×48×48×16
2	Leaky ReLU	-	-	-	48×48×48×16	48×48×48×16
3	MaxPool	-	-	2×2×2	24×24×24×16	24×24×24×16
4	Conv.	32	3×3×3	1×1×1	24×24×24×16	24×24×24×32
5	Leaky ReLU	-	-	-	24×24×24×32	24×24×24×32
6	MaxPool	-	-	2×2×2	12×12×12×32	12×12×12×32
7	Conv.	64	3×3×3	1×1×1	12×12×12×32	12×12×12×64
8	Leaky ReLU	-	-	-	12×12×12×64	12×12×12×64
9	MaxPool	-	-	2×2×2	12×12×12×64	6×6×6×64
10	Conv.	128	3×3×3	1×1×1	6×6×6×64	6×6×6×128
11	Leaky ReLU	-	-	-	6×6×6×128	6×6×6×128
12	Dropout	-	-	-	6×6×6×128	6×6×6×128
13	Transposed Conv.	64	3×3×3	2×2×2	6×6×6×128	12×12×12×64
14	Leaky ReLU	-	-	-	12×12×12×64	12×12×12×64
15	Add Layer 8	-	-	-	12×12×12×64	12×12×12×64
16	Conv.	64	3×3×3	1×1×1	12×12×12×64	12×12×12×64
17	Leaky ReLU	-	-	-	12×12×12×64	12×12×12×64
18	Transposed Conv.	32	3×3×3	2×2×2	12×12×12×64	24×24×24×32
19	Leaky ReLU	-	-	-	24×24×24×32	24×24×24×32
20	Add Layer 5	-	-	-	24×24×24×32	24×24×24×32
21	Conv.	32	3×3×3	1×1×1	24×24×24×32	24×24×24×32
22	Leaky ReLU	-	-	-	24×24×24×32	24×24×24×32
23	Transposed Conv.	16	3×3×3	2×2×2	24×24×24×32	48×48×48×16
24	Leaky ReLU	-	-	-	48×48×48×16	48×48×48×16
25	Add Layer 2	-	-	-	48×48×48×16	48×48×48×16
26	Conv.	16	3×3×3	1×1×1	48×48×48×16	48×48×48×16
27	Leaky ReLU	-	-	-	48×48×48×16	48×48×48×16
28	Conv	1	1×1×1	1×1×1	48×48×48×1	48×48×48×1
29	Leaky ReLU	-	-		48×48×48×1	48×48×48×1

S-II. EVALUATION WITH CHANNELIZED MULTI-TEMPLATE OBSERVER

A. Background

To assess the robustness of DEMIST method across various channelized model observers, we performed our objective evaluation study using a different channelized model observer, namely a channelized multi-template observer (CMTO). In our population, the defect sizes, severities and locations were all varying. In this case, it has been observed that the channel outputs (vectors) for the entire population may not be multivariate normally distributed [1], thus limiting the applicability of the widely known Hotelling observer. However, the channel outputs for sub-ensembles of patient data grouped based on defect type may have multivariate normal distributions [2]. For this case, a CMTO was developed and evaluated in the context of MPI SPECT [2]. The CMTO applies the Hotelling template to the channel outputs and adds a constant term to compute test statistics for each sub-ensemble and calculates a single global area under the ROC curve (AUC) using the pooled test statistics from all the sub-ensembles. The observer yields the maximal AUC when shifting the distributions of Hotelling observer test statistics by a different constant for each sub-ensemble is allowed [2]. The channels chosen for this observer were also the rotationally symmetric frequency channels, as in the CHO study. Also, the clinical task for this observer was detecting perfusion defects where the defect location was known.

B. Generation of test statistic

We followed the same procedure as described in Sec. III-C of the manuscript to obtain channel vectors. For each sub-ensemble, the channel vectors of defect-present and defect-absent population were used to learn the template using a leave-one-out approach. The collection of test statistic from each sub-ensemble were then pooled [2] and were used to perform the ROC analysis.

C. Results

Fig. S-1 shows the AUC values obtained with the low-dose protocol, TADL method, DEMIST method and normal-dose protocol. We observed that at all dose levels, the DEMIST method yielded a significant improvement (p < 0.05) in performance on detection task compared to the low-dose protocol as well as the TADL method. The TADL method generally did not improve performance compared to the low-dose protocol.

Figs. S-2 and S-3 show the AUC values obtained for stratified analysis based on sex. We observed that at all dose levels and stratified groups, the DEMIST method yielded a significant improvement (p < 0.05) in performance on detection task compared to the low-dose protocol and TADL method. Again, we observed that the TADL method generally did not improve performance significantly compared to the low-dose protocol.

Figs. S-4 and S-5 show the AUC values obtained for stratified analysis based on defect extent and severity, respectively. Similar to previous results, we observed that at all dose levels and stratified groups, the DEMIST method yielded a significant improvement (p < 0.05) in performance on detection task compared to low-dose protocol and TADL method. Again, the TADL method generally did not improve performance significantly compared to the low-dose protocol.

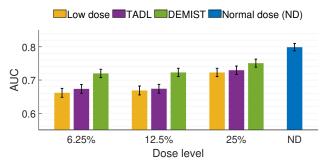


Fig. S-1. AUC values obtained for the normal and low-dose images, the images denoised using the proposed DEMIST approach and TADL approach at various dose levels with CMTO. Error bars denote 95% confidence intervals.

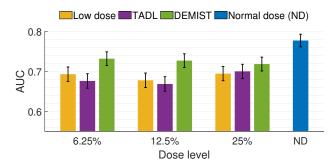


Fig. S-2. AUC values for male patients obtained for the different approaches and at various dose levels using CMTO. Error bars denote 95% confidence intervals.

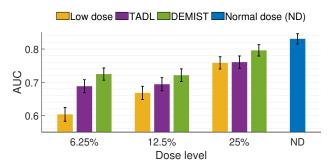


Fig. S-3. AUC values obtained for the different approaches and at various dose levels with female patients using CMTO. Error bars denote 95% confidence intervals.

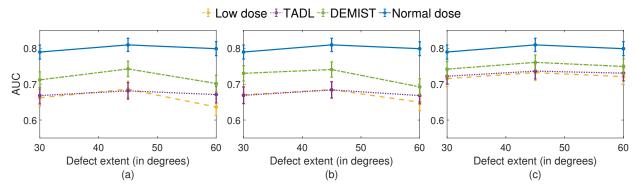


Fig. S-4. AUC values obtained using CMTO for the various approaches as a function of different defect extents with (a) 6.25%, (b) 12.5% and (c) 25% dose levels. Error bars denote 95% confidence intervals.

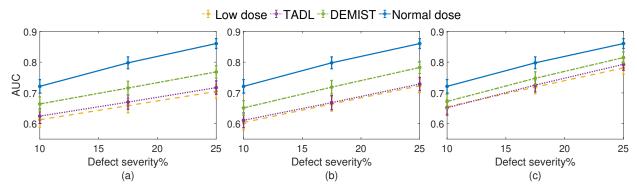


Fig. S-5. AUC values obtained using CMTO for the various approaches as a function of different defect severities with (a) 6.25%, (b) 12.5% and (c) 25% dose levels. Error bars denote 95% confidence intervals.

S-III. Corrected p-values of Delong's test for AUC comparison between two methods

TABLE S-2

CORRECTED P-VALUES FOR VARIOUS ANALYSES AND OBSERVERS (LD = LOW-DOSE PROTOCOL)

Analysis type	Stratified group	Observer	Comparisons		Dose levels	
Analysis type	Stratified group	Observer	Comparisons	6.25%	12.50%	25%
			DEMIST VS LD	2.72E-32	8.05E-38	1.44E-28
		CMTO	DEMIST VS TADL	2.74E-29	1.19E-29	2.08E-14
N			TADL VS LD	0.111834	0.840879	0.079587
Non-stratified analysis	-	СНО	DEMIST VS LD	8.78E-20	5.72E-10	2.64E-17
			DEMIST VS TADL	5.87E-11	3.71E-17	6.22E-08
			TADL VS LD	0.014445	1.634715	0.035604
			111111111111111111111111111111111111111	0.01	1100 1710	0.000000.
		СМТО	DEMIST VS LD	4.43E-07	1.83E-15	4.8E-09
	Male		DEMIST VS TADL	2.33E-18	4.39E-18	0.000553
			TADL VS LD	0.130437	0.626301	0.607878
			DEMIST VS LD	0.130437	4.71E-05	7.44E-07
		СНО	DEMIST VS TADL	8.25E-10	0.000263	0.018342
			TADL VS LD	0.130599	1.255887	0.350955
Stratified analysis based on sex			IADL VS LD	0.130399	1.233667	0.330933
Stratified aliarysis based on sex			DEMIST VS LD	2.59E-49	1.21E-16	1.13E-26
		СМТО	DEMIST VS TADL	5.46E-05	1.21E-10 1.04E-05	3.56E-17
			TADL VS LD	6.51E-15	0.001287	1.792665
	Female					
		CITO	DEMIST VS LD	1.51E-18	3.47E-07	2.46E-13
		CHO	DEMIST VS TADL	0.132264	1.27E-17	1.94E-08
	1	1	TADL VS LD	8.02E-11	0.200286	0.41967
			DEMIST VS LD	2.08E-08	4.02E-15	1.04E-09
		CMTO	DEMIST VS TADL	7.07E-10	5.29E-14	4.73E-05
	30° extent		TADL VS LD	1.581237	2.56014	0.606618
	30 CAULI		DEMIST VS LD	1.4E-05	0.025992	1.57E-06
		CHO	DEMIST VS TADL	2.42E-05	0.000472	0.00423
			TADL VS LD	1.077156	2.296377	0.362862
			DEMIST VS LD	1.1E-11	1.08E-15	9.54E-13
	45° extent	CMTO	DEMIST VS TADL	1.37E-17	4.72E-15	1.53E-06
Ct4: f - d l: - h d d - f t t t			TADL VS LD	1.904022	2.836206	1.246464
Stratified analysis based on defect extent		СНО	DEMIST VS LD	2.75E-09	2.93E-05	0.000266
			DEMIST VS TADL	5.04E-07	4.47E-08	0.004428
			TADL VS LD	0.681075	1.96272	2.055168
			DEMIST VS LD	1.31E-13	4.45E-08	4.55E-08
		СМТО	DEMIST VS TADL	7.56E-05	0.002286	0.000327
	600		TADL VS LD	0.002466	0.130788	0.251469
	60° extent	СНО	DEMIST VS LD	1.08E-10	2.12E-06	2.58E-08
			DEMIST VS TADL	0.088839	3.15E-07	0.001602
			TADL VS LD	0.000753	1.500372	0.080946
			1			
	10% severity	СМТО	DEMIST VS LD	1.11E-07	1.21E-07	0.001269
			DEMIST VS TADL	4.07E-07	6.94E-06	0.001209
			TADL VS LD	0.784584	1.289538	1.785861
			DEMIST VS LD	0.764364	0.038628	0.001143
			DEMIST VS TADL	0.000343	0.038028	0.001143
			TADL VS LD	0.142011	2.395917	0.808731
			INDE VOED	0.540419	2.373711	0.000731
	17.5% severity	СМТО	DEMIST VS LD	2.88E-11	1.56E-13	2.73E-10
			DEMIST VS LD DEMIST VS TADL		5.92E-11	
				2.83E-10		9.09E-06
Stratified analysis based on defect severity			TADL VS LD	0.719325	1.793304	0.692793
		СНО	DEMIST VS LD	2.35E-07	0.000505	9.99E-07
			DEMIST VS TADL	0.00026	1.07E-06	0.001629
			TADL VS LD	0.312813	2.061603	0.514611
		G) fmc	DEMICE VC IP	7.045.16	5 215 21	1 225 17
			DEMIST VS LD	7.94E-16	5.31E-21	1.33E-17
		a		201=::		
		СМТО	DEMIST VS TADL	3.04E-14	1.6E-15	3.68E-07
	25% severity	СМТО	DEMIST VS TADL TADL VS LD	0.542664	1.39266	0.038385
	25% severity		DEMIST VS TADL TADL VS LD DEMIST VS LD	0.542664 1.58E-12	1.39266 4.38E-07	0.038385 6.98E-10
	25% severity	СМТО	DEMIST VS TADL TADL VS LD	0.542664	1.39266	0.038385

S-IV. SPECT SCANNER CONFIGURATION

TABLE S-3

ACQUISITION AND RECONSTRUCTION PARAMETERS OF SPECT/CT SYSTEMS. (LEHR=LOW-ENERGY HIGH-RESOLUTION, WEHR=WIDE-ENERGY HIGH-RESOLUTION)

	Scanner				
	GE Discovery NM/CT 670 Pro NaI	GE Discovery NM/CT 670 Pro CZT			
Number of cases in test data	63	51			
Number of cases in train data	102	82			
Number of cases in validation data	12	28			
Collimator type	LEHR	WEHR			
Collimator grid	Parallel hole	Parallel hole			
Detector	NaI	CZT			
Energy resolution at 140 keV (%)	9.8	6.3			
Intrinsic spatial resolution (in mm)	3.9	2.46			
System sensitivity (cps/MBq) at 10 cm	72	85			
Photopeak energy window (in keV)	126-154	126-154			
Reconstruction	OSEM	OSEM			
Subsets	6	6			
Iteration	8	8			
Attenuation correction	CT	CT			

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 X. Li, A. K. Jha, M. Ghaly, F. E. Elshahaby, J. M. Links, and E. C. Frey, "Use of sub-ensembles and multi-template observers to evaluate detection task performance for data that are not multivariate normal," *IEEE Trans. Med. Imaging*, vol. 36, no. 4, pp. 917–929, 2016.