12th Annual Otolaryngology Literature Update Sinus & Rhinology I

Rodney J. Schlosser, M.D.

Professor

Director, Rhinology & Sinus Surgery
Department of Otolaryngology - Head & Neck Surgery
Medical University of South Carolina
schlossr@musc.edu

Rodney J. Schlosser, M.D. is the Director of Rhinology and Sinus Surgery at MUSC. After graduating from the U.S. Military Academy at West Point and serving four years of active military duty in the U.S. Army, Dr. Schlosser received his medical degree from the Mayo Clinic. He then completed his ENT residency at the University of Virginia followed by an additional year of fellowship devoted exclusively to the treatment of advanced sinus disease at the University of Pennsylvania with David W. Kennedy, M.D. one of the premiere sinus surgeons in the world.

Dr. Schlosser has written a textbook on the management of sinus problems and more than 350 articles and chapters dealing with sinus and nasal topics. He regularly travels around the world to speak and teach courses on the latest techniques in endoscopic sinus surgery and computerized image-guided surgery. Dr. Schlosser focuses primarily on difficult adult and pediatric sinus cases, nasal obstruction, revision surgeries, sinonasal tumors.

.

12th Annual Otolaryngology Literature Update Medical University of South Carolina

Sinus & Rhinology I Rodney J. Schlosser, M.D.

- Bachert C, Corren J, Lee SE, Zhang H, Harel S, Cunoosamy D, Khan AH, Jacob-Nara JA, Siddiqui S, Nash S, Rowe PJ, Deniz Y. Dupilumab efficacy and biomarkers in chronic rhinosinusitis with nasal polyps: Association between dupilumab treatment effect on nasal polyp score and biomarkers of type 2 inflammation in patients with chronic rhinosinusitis with nasal polyps in the phase 3 SINUS-24 and SINUS-52 trials. Int Forum Allergy Rhinol. 2022 Sep;12(9):1191-1195. doi: 10.1002/alr.22964. Epub 2022 Jan 31. PMID: 34970860; PMCID: PMC9544911.
- Boscolo-Rizzo P, Hummel T, Invitto S, Spinato G, Tomasoni M, Emanuelli E, Tofanelli M, Cavicchia A, Grill V, Vaira LA, Lechien JR, Borsetto D, Polesel J, Dibattista M, Menini A, Hopkins C, Tirelli G. Psychophysical assessment of olfactory and gustatory function in post-mild COVID-19 patients: A matched case-control study with 2-year follow-up. Int Forum Allergy Rhinol. 2023 Feb 28. doi: 10.1002/alr.23148. Epub ahead of print. PMID: 36852674.
- Boscolo-Rizzo P, Tirelli G, Meloni P, Hopkins C, Madeddu G, De Vito A, Gardenal N, Valentinotti R, Tofanelli M, Borsetto D, Lechien JR, Polesel J, De Riu G, Vaira LA. Coronavirus disease 2019 (COVID-19)-related smell and taste impairment with widespread diffusion of severe acute respiratory syndrome-coronavirus-2 (SARS-CoV-2) Omicron variant. Int Forum Allergy Rhinol. 2022 Oct;12(10):1273-1281. doi: 10.1002/alr.22995. Epub 2022 Mar 24. PMID: 35286777; PMCID: PMC9082058.
- Dehgani-Mobaraki P, Patel Z, Zaidi AK, Giannandrea D, Hopkins C. The Omicron variant of SARS-CoV-2 and its effect on the olfactory system. Int Forum Allergy Rhinol. 2023 May;13(5):958-960. doi: 10.1002/alr.23089. Epub 2022 Sep 29. PMID: 36103157; PMCID: PMC9539397.
- Garcia JAP, Miller E, Norwood TG, Dorin NA, Grayson J, Woodworth B, Cho DY. Gabapentin improves parosmia after COVID-19 infection. Int Forum Allergy Rhinol. 2023 Jun;13(6):1034-1036. doi: 10.1002/alr.23117. Epub 2022 Dec 15. PMID: 36484400; PMCID: PMC9877759.
- Lee DJ, Cramer HB, Kshirsagar RS, Douglas JE, Kohanski MA, Palmer JN, Adappa ND, Bosso JV. Dupilumab-related adverse events among patients with chronic rhinosinusitis with nasal polyposis. Int Forum Allergy Rhinol. 2022 Nov 17. doi: 10.1002/alr.23108. Epub ahead of print. PMID: 36394542.
- Miglani A, Soler ZM, Smith TL, Mace JC, Schlosser RJ. A comparative analysis of endoscopic sinus surgery versus biologics for treatment of chronic rhinosinusitis with nasal polyposis. Int Forum Allergy Rhinol. 2023 Feb;13(2):116-128. doi: 10.1002/alr.23059. Epub 2022 Sep 4. PMID: 35980852; PMCID: PMC9877092.
- Yan CH, Jang SS, Lin HC, Ma Y, Khanwalkar AR, Thai A, Patel ZM. Use of platelet-rich plasma for COVID-19-related olfactory loss: a randomized controlled trial. Int Forum Allergy Rhinol. 2023 Jun;13(6):989-997. doi: 10.1002/alr.23116. Epub 2022 Dec 21. PMID: 36507615; PMCID: PMC9877663.

Literature course 2023

Rodney J. Schlosser, MD



Professor and Director of Rhinology Department of Otolaryngology - HNS Medical University of South Carolina Charleston, SC



Two hot topics

Biologics & COVID-anosmia





Dupilumab – Death of a profession?

Rodney J. Schlosser, MD



Professor and Director of Rhinology Department of Otolaryngology - HNS Medical University of South Carolina Charleston, SC



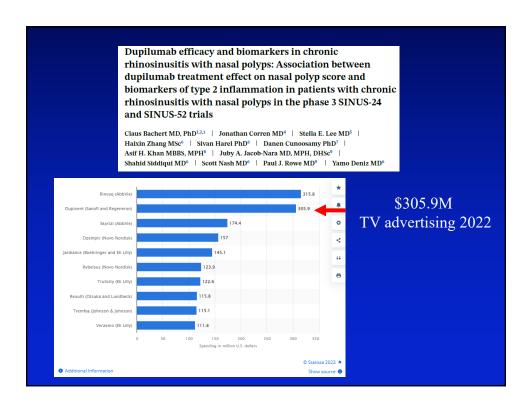
Disclosures

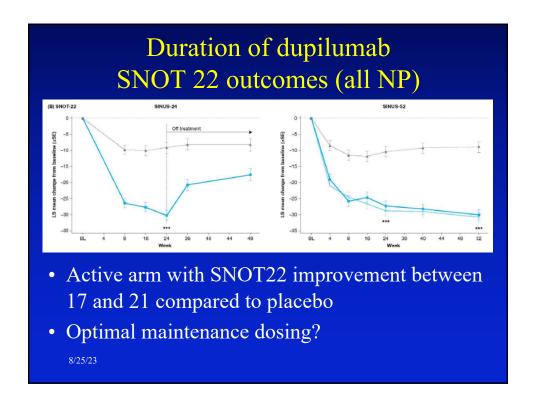
• Grant support/consulting: Sanofi, GSK, Genentech, Xhance, Cyrano, Sinusonic

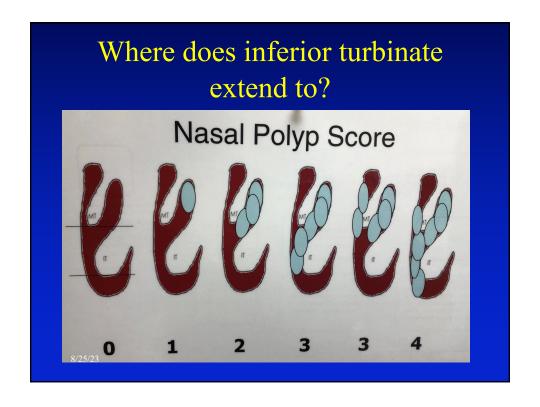
What do we do with this patient?

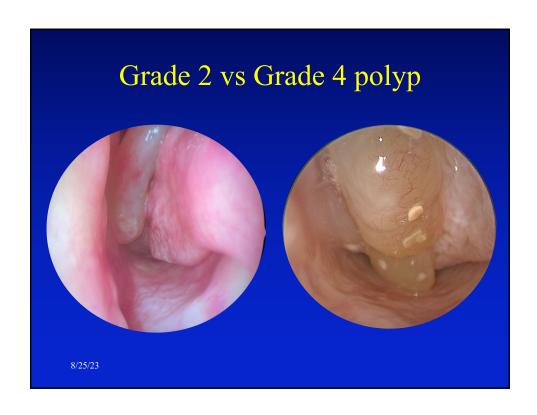


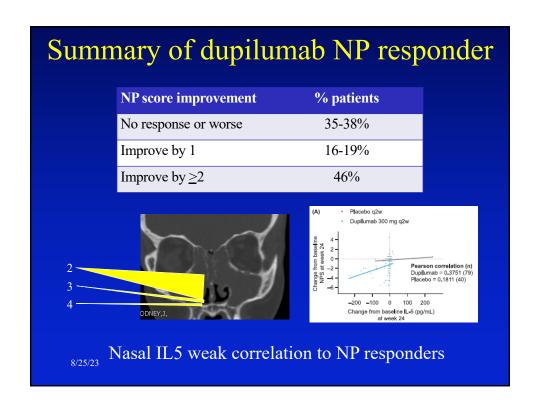
Surgery first, then biologic for ESS failure?
Which surgery? Lothrop?
Biologics first, ESS for biologic failure?
Any way to predict treatment response?











How does ESS compare?

- "Polyps always recur"
- "You will need surgery annually for rest of your life"
- JUST THE FACTS NO SPIN!

8/25/23

Sino-Nasal Outcome Test-22 Outcomes After Sinus Surgery: A Systematic Review and Meta-Analysis

Zachary M. Soler, MD, MSc ©; Rabun Jones, BS; Phong Le, MD; Luke Rudmik, MD; Jose L. Mattos, MD, MPH; Shaun A. Nguyen, MD ©; Rodney J. Schlosser, MD

Systematic Review and Meta-analysis of SNOT-22 Outcomes after Surgery for Chronic Rhinosinusitis with Nasal Polyposis

Phong T. Le, MD¹, Zachary M. Soler, MD¹, Rabun Jones¹, Jose L. Mattos, MD^{1,2}, Shaun A. Nguyen, MD¹, and Rodney J. Schlosser, MD¹

- SR/MA show mean SNOT22 improvement 23-24 pts
- Greater SNOT22 improvements after ESS in patients with asthma regardless of NP status, preop SNOT22

CRSwNP revision rates: Meta-analysis Starting to see impact of phenotypes

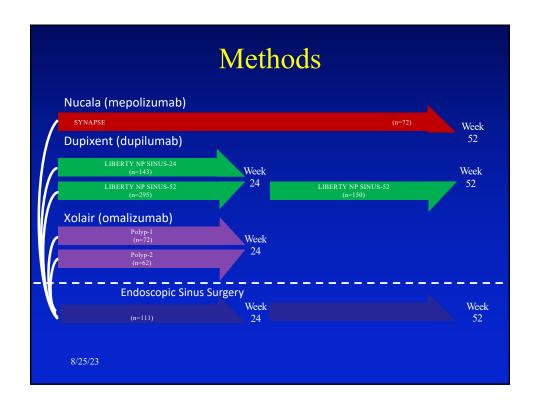
Subgroup	Total # of patients (n)	% Revision	% Difference	95% CI	p value
All Studies	34,220	18.6%	8.5	4.6-12.7	p<0.0001
AERD	465	27.2%	8.3	4.0-12.7	p<0.0001
All Studies	34,220	18.6%	10.1	6.1-14.4	p<0.0001
AFRS	467	28.7%	10.1	0.1-14.4	p<0.0001
AERD	465	27.2%	1.6	-4.2-7.3	p=0.5902
AFRS	467	28.7%	1.0	-4.2-7.3	p=0.3902
Asthma	552	22.6%	14.6	5.6-20.3	p=0.0035
No asthma	75	8.0%	14.0	5.0-20.5	p=0.0033
Primary ESS	10030	14.3%	12.1	11 0 12 2	·· <0.0001
Any Revision ESS	9910	26.4%	12.1	11.0-13.2	p<0.0001
Studies prior to 2008	2053	22.7%	6.1	4.3-8.0	p<0.0001
Studies after 2008	32264	16.6%			•

8/25/23

Loftus C, etal, IFAR 2020

Prospective, randomized, blinded trial of biologic vs ESS will never be done.....BUT

A comparative analysis of endoscopic sinus surgery versus biologics for treatment of chronic rhinosinusitis with nasal polyposis



Outcomes • PROMs - SNOT22 - Nasal Congestion Score (0-3) - Smell Score (0-3) • Nasal Polyp Score (endoscopy) • Smell tests: SIT40 or Sniffin Sticks

	Endpoints	Baseline (mean <u>+</u> SD)	Week 24 (mean±SD)	LS mean change from baseline (±SE)	
ESS	NCS	2.9 [±0.3]	0.9 [±0.9]	-1.9 [±0.09]	
	Loss of smell				
(n=111)	score	2.7 [±0.7]	1.5 [±1.2]	-1.2 [±0.13]	
	SNOT 22 score	56.1 [±19.6]	22.9 [±19.6]	-33.3 [±1.8]	Statistics
		Baseline	Week 24	LS mean change from	(comparison of
	Endpoints	(mean+SD)	(mean+SD)	baseline (±SE)	mean Δ to ESS)
Dupi-24	NCS	2.26 [±0.57]	0.94 [±0.75]	-1.34 [+0.07]	p<0.001
	Loss of smell			210 1 (210 1)	
(n=143)	score	$2.70 [\pm 0.57]$	1.35 [±0.99]	-1.41 [<u>+</u> 0.07]	NS
	SNOT 22 score	48.00 [±20.16]	18.58 [±14.92]	-30.43 [±1.54]	NS
		Baseline	Week 24	LS mean change from	Statistics (comparison of
	Endpoints	(mean±SD)	(mean±SD)	haseline (±SE)	mean Δ to ESS)
Dupi-52	NCS	2.46 [±0.77]	1.19 [±0.90]	-1.25[±0.06]	p<0.001
	Loss of smell				
(n=295)	score	$2.77 [\pm 0.77]$	1.55 [±1.02]	-1.21[±0.06]	NS
	SNOT 22 score	51.0 [±4.67]	23.89 [±18.77]	-27.77[±1.26]	p=0.018

	Endpoints	Baseline (mean <u>+</u> SD)	Week 24 (mean±SD)	LS mean change from baseline (±SE)	
ESS	NCS	2.9 [±0.3]	0.9 [±0.9]	-1.9 [±0.09]	
(n=111)	Loss of smell score SNOT 22 score	2.7 [±0.7] 56.1 [±19.6]	1.5 [±1.2] 22.9 [±19.6]	-1.2 [±0.13] -33.3 [±1.8]	
	Endpoints	Baseline (mean+SD)	Week 24 (mean+SD)	LS mean change from baseline (±SE)	Statistics (comparison of mean Δ to ESS)
Dupi-24	NCS	2.26 [±0.57]	0.94 [±0.75]	-1.34 [<u>+</u> 0.07]	p<0.001
(n=143)	Loss of smell score SNOT 22 score	2.70 [±0.57] 48.00 [±20.16]	1.35 [±0.99] 18.58 [±14.92]	-1.41 [<u>+</u> 0.07] -30.43 [+1.54]	NS NS
	Endpoints	Baseline (mean+SD)	Week 24 (mean+SD)	LS mean change from baseline (±SE)	Statistics (comparison of mean A to ESS)
Dupi-52	NCS	2.46 [±0.77]	1.19 [±0.90]	-1.25[±0.06]	p<0.001
(n=295)	Loss of smell score SNOT 22 score	2.77 [±0.77] 51.0 [±4.67]	1.55 [±1.02] 23.89 [±18.77]	-1.21[±0.06] 27.77[±1.26]	NS p=0.018

				LS mean change from	
	Endpoints	(mean+SD)	(mean+SD)	hasolino (+SE)	
ESS	NCS	2.9 [±0.3]	0.9 [±0.9]	-1.9 [±0.09]	
	Loss of smell				
(n=111)	score	2.7 [±0.7]	1.5 [±1.2]	-1.2 [±0.13]	
	SNOT 22 score	56.1 [±19.6]	22.9 [±19.6]	-33.3 [±1.8]	
			Week 24 (mean)		Statistics
		Baseline	(Unable to	LS mean change from	(comparison of
	Endpoints	(mean <u>+</u> SD)	calculate SD)	baseline (±SE)	mean Δ to ESS)
Oma-1	NCS	$2.4 [\pm 0.7]$	1.51	0.89 [±0.10	p<0.001
	Loss of smell				
(n=72)	score	$2.5 [\pm 0.8]$	1.94	-0.56 [±0.09]	P<0.001
	SNOT 22 score	59.8 [±19.7]	35.1	-24.70 [±2.01]	p=0.002
			Week 24 (mean)		Statistics
		Baseline	(Unable to	LS mean change from	(comparison of
	Endpoints	(mean+SD)	calculate SD)	baseline (±SE)	mean Δ to ESS)
Oma-2	NCS	$2.3 [\pm 0.7]$	1.6	-0.70 [±0.11	p<0.001
	Loss of smell				
(n=62)	score	2.6 [±0.8]	2.02	-0.58 [±0.10]	p=0.001
	SNOT 22 score	59.2 [±20.5]	37.61	-21.59 [±2.25]	p<0.001

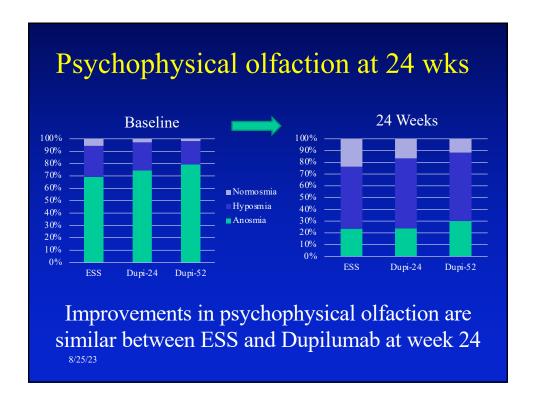
		Baseline	Week 24	LS mean change from	
	Endpoints	(mean <u>+</u> SD)	(mean±SD)	baseline (±SE)	
ESS	NCS	2.9 [±0.3]	0.9 [±0.9]	-1.9 [±0.09]	
	Loss of smell				
(n=111)	score	2.7 [±0.7]	1.5 [±1.2]	-1.2 [±0.13]	
	SNOT 22 score	56.1 [±19.6]	22.9 [±19.6]	-33.3 [±1.8]	Or at at
			Week 24 (mean)		Statistics
		Baseline	(Unable to	LS mean change from	(comparison of
0 1	Endpoints	(mean+SD)	calculate SD)	baseline (±SE)	mean Δ to ESS)
Oma-1	NCS Loss of smell	2.4 [±0.7]	1.51	-0.89 [±0.10]	p<0.001
(-7 2)		2.5 [0.0]	1.94	-0.56 [±0.09]	P<0.001
(n=72)	score SNOT 22 score	2.5 [±0.8] 59.8 [±19.7]	35.1	-0.56 [±0.09] -24.70 [±2.01]	p=0.002
	SNOT 22 score	39.6 [±19.7]	Week 24 (mean)	24.70 [±2.01]	Statistics
		Baseline	(Unable to	LS mean change from	(comparison of
	Endpoints	(mean+SD)	calculate SD)	baseline (±SE)	mean Δ to ESS)
Oma-2	NCS	2.3 [±0.7]	1.6	-0.70 [±0.11]	p<0.001
	Loss of smell	[]	***	01.0 [-0121]	P 0.001
(n=62)	score	2.6 [±0.8]	2.02	-0.58 [±0.10]	p=0.001
	SNOT 22 score	59.2 [±20.5]	37.61	21.59 [±2.25]	p<0.001
8/25/2	cvn			SS offers s 24 weeks	uperior

	Endonina	Baseline	Week 52	LS mean from baseline (±SE)	
ESS	Endpoints NCS (n=48)	(mean±SD) 2.9 [±0.3]	(mean+SD) 0.8 [±0.8]	-2.1 [±0.13]	
200	Loss of smell	200 [=000]	0.0 [=0.0]	211 (=0110)	
	score (n=48)	$2.8 [\pm 0.5]$	1.6 [±1.2]	-1.2 [±0.16]	
	SNOT-22	•			
	score (n=48)	57.7 [±18.9]	23.8 [±19.1]	-33.9 [±3.04]	
					Statistics
		Baseline	Week 52	LS mean from	(compariso
	Endpoints	(mean±SD)	(mean+SD)	baseline (±SE)	n to ESS)
Dupi-52 (group A					
q2w dosing)	NCS	$2.48 [\pm 0.62]$	1.10 [±0.92]	1.35[+0.07]	p<0.001
	Loss of smell				
(n=150)	score	2.81 [±0.46]			
	SNOT 22		21.67		***
	score	50.16 [±19.72]	[±19.16]	-29.84[+1.63]	NS

Loss of smell score (n=48) 2.8 [±0.5] 1.6 [±1.2] -1.2 [±0.16] SNOT-22 score (n=48) 57.7 [±18.9] 23.8 [±19.1] 33.9 [±3.04]	
SNOT-22 score (n=48) 57.7 [±18.9] 23.8 [±19.1] 33.9 [±3.04]	
score (n=48) 57.7 [±18.9] 23.8 [±19.1] 33.9 [±3.04]	
DE W1-52 I C (Statistics
	compariso
	n to ESS)
Dupi-52 (group A	
q2w dosing) NCS 2.48 $[\pm 0.62]$ 1.10 $[\pm 0.92]$ -1.35 $[\pm 0.07]$	p<0.001
Loss of smell (n=150) score 2.81 [±0.46]	
SNOT 22 21.67	
score 50.16 [±19.72] [±19.16] -29.84[+1.63]	NS

	Endpoints	Baseline (mean+SD)	Week 52 (mean+SD)	LS mean from baseline (±SE)	
ESS	NCS (n=48)	2.9 [±0.3]	0.8 [±0.8]	-2.1 [±0.13]	
	Loss of smell		, ,		
	score (n=48) SNOT-22 score (n=48)	2.8 [±0.5] 57.7 [±18.9]	1.6 [±1.2] 23.8 [±19.1]	-1.2 [±0.16] -33.9 [±3.04]	
	Endpoints	Baseline (mean+SD)	Week 52 (mean+SD)	LS mean from	Statistics (comparison to ESS)
Меро	NCS	2.67 [±0.24]	1.4	-1.26 [<u>+</u> 1.03]	p<0.001
n=206)	Loss of smell score	2.88 [<u>+</u> 0.24]	2	-0.84[<u>+</u> 1.08]	p=0.039
	SNOT 22 score	63.7 [±17.6]	34.3	-29.4 [±24.67]	NS
Con	nared to	Menolis	zumah i	ESS offe	rs similar

	Endpoints	Baseline (mean+SD)	Week 52 (mean+SD)	LS mean from baseline (±SE)	
ESS	Dira points	(mean is b)	(mean · SD)	baseriie (±32)	
	NCS (n=48)	2.9 [±0.3]	0.8 [±0.8]	-2.1 [±0.13]	
	Loss of smell score (n=48)	2.8 [±0.5]	1.6 [±1.2]	-1.2 [±0.16]	
	SNOT-22 score (n=48)	57.7 [±18.9]	23.8 [±19.1]	-33.9 [±3.04]	
	Endpoints	Baseline (mean+SD)	Week 52 (mean+SD)	LS mean from baseline (±SE)	Statistics (comparison to ESS)
Меро	NCS	2.67 [+0.24]	1.4	-1.26 [+1.03]	p<0.001
n=206)	Loss of smell score	2.88 [±0.24]	2	-0.84[<u>+</u> 1.08]	p=0.039
	SNOT 22 score	63.7 [±17.6]	34.3	29.4 [±24.67]	NS
Con	npared to	Mepoliz	zumab, I	ESS offe	rs similar



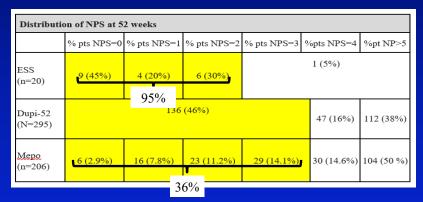
Lund	l Kennedy Polyp Score (LK-NP)		Nasal Polyp Score (NPS)
LK 0	No polyps	NPS 0	No polyps
LK 1	Polyps confined/restricted to the middle meatus	NPS 1	Polyps not reaching below inferior border of middle turbinate/confined to middle meatus
LK 2	Polyps extending beyond the middle	NPS 2	Polyps reaching below the lower border of the middle turbinate
	meatus	NPS 3	Large polyps reaching the lower border of the inferior turbinate
		NPS 4	Large polyps causing complete obstruction of the inferior nasal cavity

24-week nasal polyp scores

Distribution	of NPS at 24	weeks				
	% pts NPS=0	% pts NPS=1	% pts NPS=2	% pts NPS=3	%pt NPS=4	%pts NP>5
ESS (n=79)	48 (61%)	7 (9%)	14 (18%)		10 (13%)	
Dupi-24 (n=143)		87%	16%)		27 (19%)	50 (35%)
Oma-1&2 (n=128)		42 (3	31%)		30 (25%)	56 (44 %)

ESS results in significantly lower NPS compared to Dupilumab and Omalizumab at week 24

52-week nasal polyp scores



ESS results in significantly lower NPS compared to Dupilumab and Mepolizumab at week 52

Dupilumab-related adverse events among patients with chronic rhinosinusitis with nasal polyposis

Daniel J. Lee $MD^1 \odot \mid Hannah B. Cramer BS^1 \mid Rijul S. Kshirsagar <math>MD^2 \odot \mid Jennifer E. Douglas MD^1 \odot \mid Michael A. Kohanski MD, PhD^1 \mid James N. Palmer MD^1 \mid Nithin D. Adappa MD^1 \mid John V. Bosso MD^1$

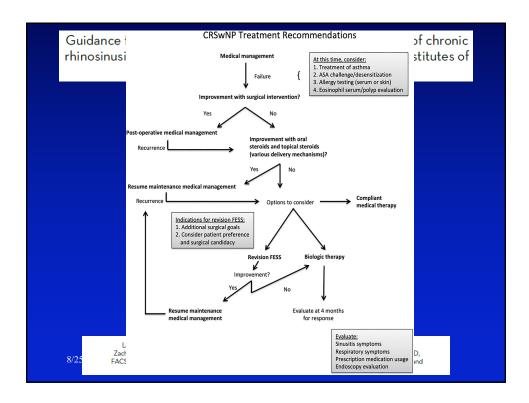
- N=58 consecutive pts
- 24.1% with AEs requiring d/c of dupi
- Rash 42.9%; joint pain 36%

8/25/23

Guidance for contemporary use of biologics in management of chronic rhinosinusitis with nasal polyps: discussion from a National Institutes of Health-sponsored workshop

8/25

Lauren T. Roland, MD, MSCl¹, Timothy L. Smith, MD, MPH, FACS², Rodney J. Schlosser, MD³, Zachary M. Soler, MD, MSc³, Anju T. Peters, MD, MSCl⁴, Tanya M. Laidlaw, MD³, Benjamin S. Bleier, MD, FACS⁶, Vijay Ramakrishnan, MD⁷, B. Elina Toskala, MD, PhD, MBA⁸, David W. Kennedy, MD, FACS⁹ and Amber U. Luong, MD, PhD, FACS¹⁰



Summary: Biologics and ESS

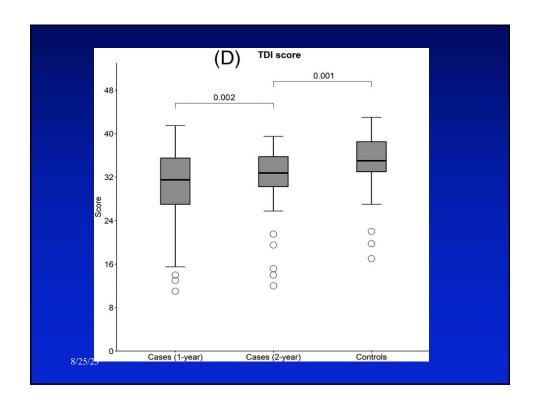
- Steroid rinses/Xhance may provide benefit prior to ESS
- ESS with postop topical steroid has good success rate
- Biologics generally reserved for ESS failures
- Hopefully have predictors in future
 - Mucus biomarkers, eosinophil count
 - Phenotype: AERD, AFRS

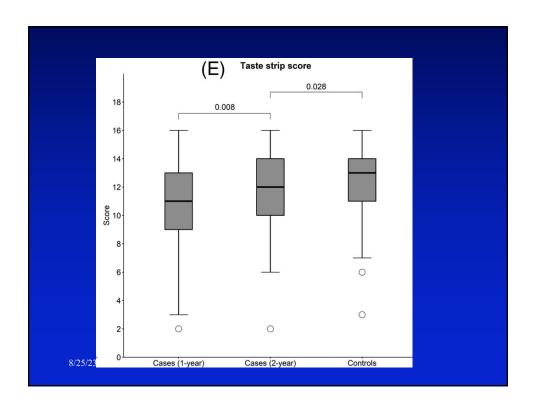
Shifting gears to COVID

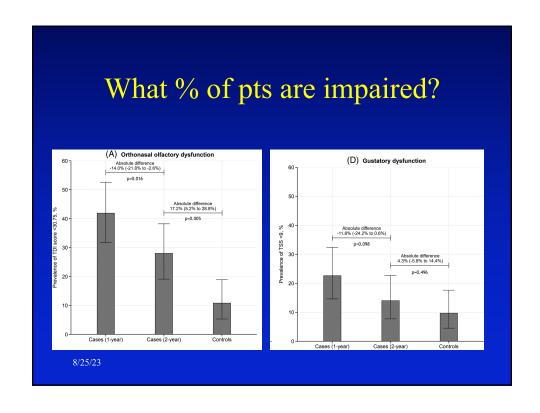
8/25/23

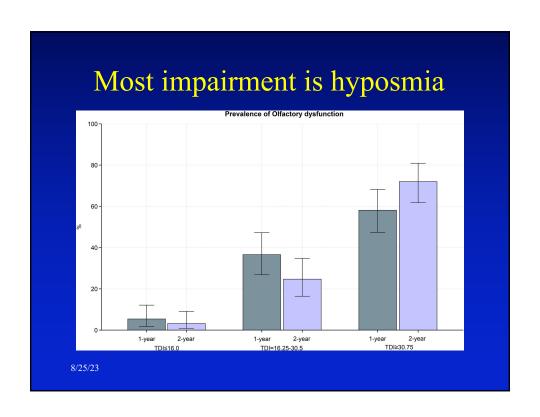
Psychophysical assessment of olfactory and gustatory function in post-mild COVID-19 patients: A matched case-control study with 2-year follow-up

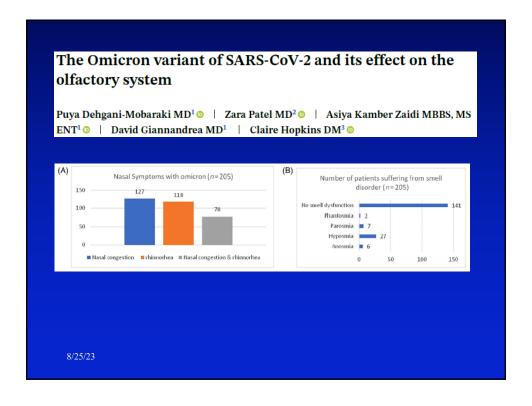
- N=93 COVID+ and 93 matched controls
- Median age 49 yrs, 59% female
- 1 and 2 year f/u post-COVID

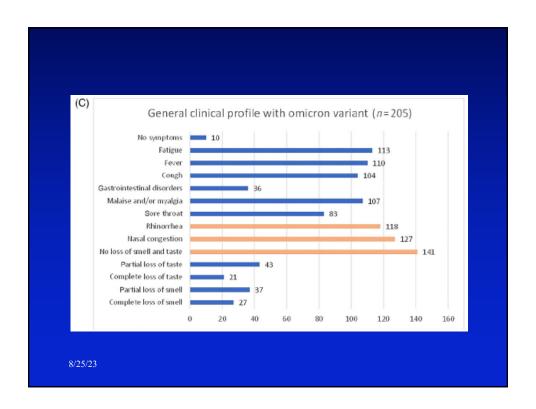












Gabapentin improves parosmia after COVID-19 infection

Jaime A. Pena Garcia $MD^1 \mid Emily \, Miller \, MD^1 \mid Timothy \, G. \, Norwood \, MD^1 \mid Natalie A. \, Dorin \, MD^1 \mid Jessica \, Grayson \, MD^1 \mid Bradford \, Woodworth \, MD^{1,2} \mid Do-Yeon \, Cho \, MD^{1,2,3} \, \bigcirc$

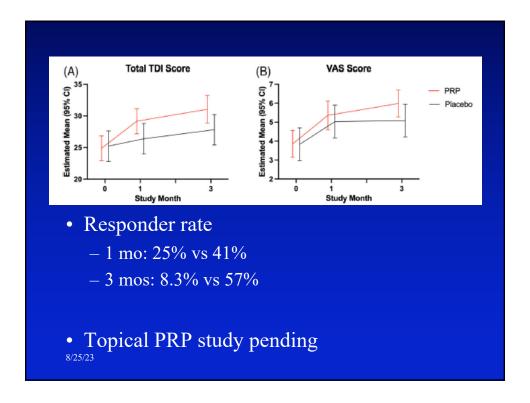
- N=12 pts, OD 7-15 mos duration
- 9 pts with phone f/u
- 100 mg qd x 7d; increase 100-300 mg weekly as tolerated
- 6/9 with significant subjective improvement
- SIT40 in 3 pts, no change

8/25/23

Use of platelet-rich plasma for COVID-19-related olfactory loss: a randomized controlled trial

Carol H. Yan MD¹ | Sophie S. Jang MD¹ | Hung-Fu C. Lin NP-C² | Yifei Ma MA, MS² | Ashoke R. Khanwalkar MD² | Anthony Thai MD² | Zara M. Patel MD² |

- N=35 pts (26 completed study), COVID-OD
- OD duration 6-12 mos w SIT40<33
- 3 injections into olfactory cleft at weeks 0,2,4 of PRP or saline
- Follow up at 1 and 3 mos



COVID dysosmia summary

- Incidence decreasing, but still a problem
- Rxs: Smell training (fifthsense.org.uk), topical steroid
- 2nd level rx: PRP, gabapentin
- Future rx: Theophylline spray?