



CLINICAL ASPECT OF TRANSPLANTS

Name: Helle Kiellberg Larsen

Credentials: Consultant, MD, PhD, associate professor

Institution/Location: Department of Dermato-venereology, Bispebjerg Hospital, Copenhagen, Denmark

DISCLOSURES

• Advisory board meeting 19.03.2024 MSD Denmark

LEARNING OBJECTIVES

Organ transplant recipients, here represented by kidney transplant recipients, from a Danish perspective, and their risk of:

Anal hrHPV infection and for female KTRs cervical co-infection

Anal warts

Anal HSIL

Perspectives for anal cancer screening in this population

BACKGROUND

IN DENMARK THERE IS NO SCREENING PROGRAM FOR ANAL CANCER, AND THIS IS REFLECTED IN LOWER RATES OF ANAL HSIL AND HIGHER RATES OF ANAL CANCER

1998 - 2018:

2580 anal cancer (90% SCC)

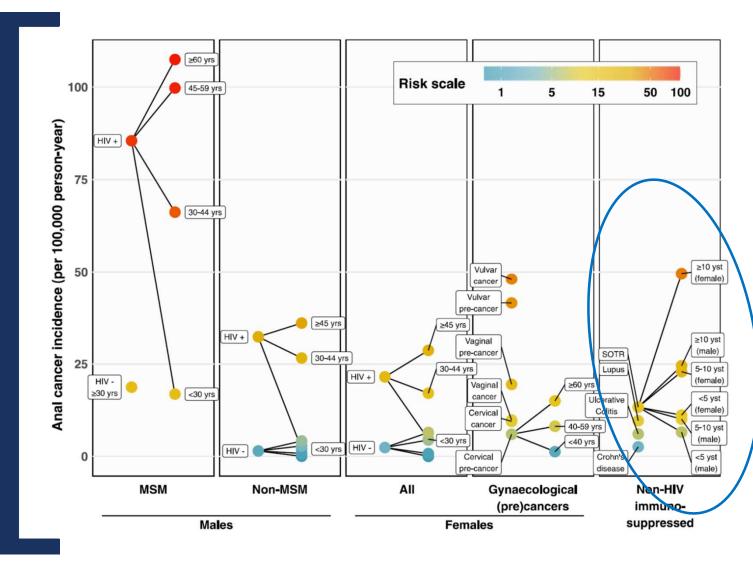
871 aHSIL-AIN2/3

2/3 of anal SCC and aHSIL-AIN2/3 were among women

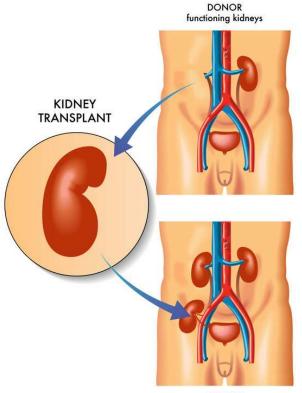
AIN: Anal intraepithelial neoplasia aHSIL: anal high grade squamous intraepithelial lesion

Ref.: Urbute A et al. Trends in incidence and survival from anal cancer and incidence of high-grade anal intraepithelial neoplasia in Denmark. Cancer Epidemiol. 2022 Apr;77:102099

REF.:
CLIFFORD G ET AL. A META-ANALYSIS OF ANAL CANCER INCIDENCE BY RISK GROUP: TOWARD A UNIFIED ANAL CANCER RISK SCALE. INT J CANCER. 2021;148(1):38-47.



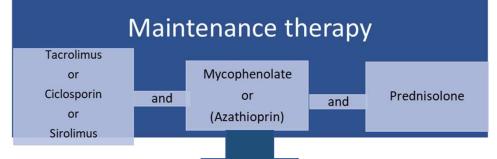
BACKGROUND







Induction therapy Interleukin II inhibitor or T-lymphocyte-depleting agent



Impaired host T-cell response on the clearance or control of an established HPV infection

	Project	overview
Registry-based study	Kidney transplar	Clinical study
Risk of genital warts		Prevalence and risk of anogenital warts
Risk of cervical-, vulvar-, vaginal-, penile- and anal precancerous lesions and cancer		Prevalence and risk of anal HPV infection Prevalence and risk of anal HSIL
		Prevalence of cervical HPV infection and risk of anal co- infektion
		Incidence and clearance of cervical and anal HPV infection
		Prevalence of oral HPV infection
		Prevalence of penile HPV infection

Registry-based study

Risk of genital warts

Risk of cervical-, vulvar-, vaginal-, penileand anal precancerous lesions and cancer

Project overview

Clinical study

Kidney transplant recipients

Prevalence and risk of anogenital warts

Prevalence and risk of anal HPV infection

Prevalence and risk of anal HSIL

Prevalence of cervical HPV infection and risk of anal coinfektion Revised: 8 July 2018 | Accepted: 25 July 2018

DOI: 10.1111/ajt.15056

ORIGINAL ARTICLE

AJT

Risk of genital warts in renal transplant recipients—A registrybased, prospective cohort study

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Helle Kiellberg Larsen<sup>1,2</sup> | Louise T. Thomsen<sup>1</sup> | Merete Haedersdal<sup>2</sup>
                                                                                                     Christian
Dehlendorff<sup>3</sup> | Søren Schwartz Sørensen<sup>4</sup> | Susanne K. Kjaer<sup>1,5</sup>
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Received: 10 August 2018

Accepted: 23 August 2018

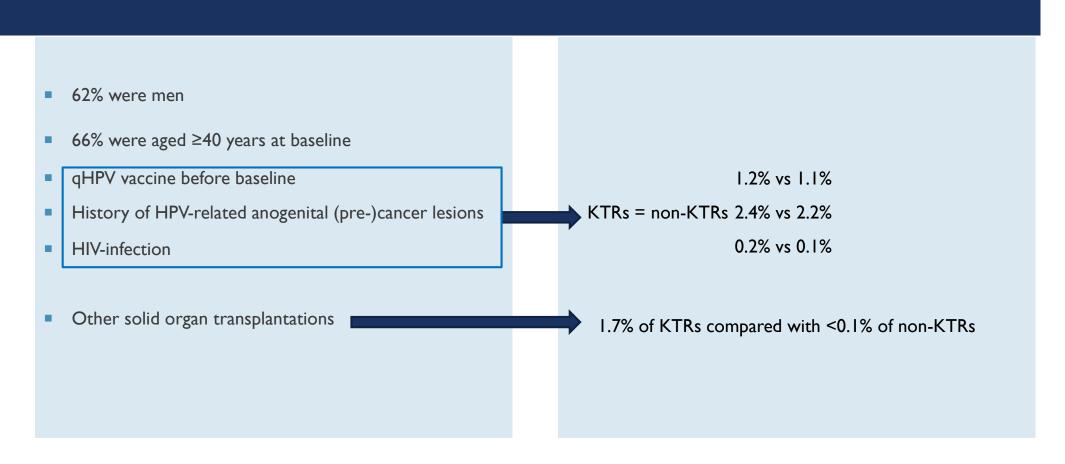
DOI: 10.1111/ajt.15108

EDITORIAL

AJT

Time to take on HPV in transplant clinic—Warts and all

RESULTS (REGISTRY-BASED STUDY) LARSEN HK ET AL. AMERICAN JOURNAL OF TRANSPLANTATION. 2019;19(1):156-165



DOI: 10.1111/ajt.15056

ORIGINAL ARTICLE

AJT

Risk of genital warts in renal transplant recipients—A registry-based, prospective cohort study

Helle Kiellberg Larsen^{1,2} | Louise T. Thomsen¹ | Merete Haedersdal² | Christian Dehlendorff³ | Søren Schwartz Sørensen⁴ | Susanne K. Kjaer^{1,5}

TABLE 2 Hazard ratio of genital warts in Danish renal transplant recipients (RTRs) 1996-2015 vs a nontransplanted control cohort (non-RTRs), overall and stratified by sex

	Person-y at risk	Number of GW episodes	HRª	95% CI	HR ^b	95% CI
Both sexes						
Non-RTRs	1 377 972	2025	1	Ref	1	Ref
RTRs	23 926	132	3.41	2.86-4.06	3.30	2.76-3.93
Women						
Non-RTRs	522 262	654	1	Ref	1	Ref
RTRs	9043	60	4.94	(3.79-6.43)	4.70	3.61-6.13
Men						
Non-RTRs	855 711	1371	1	Ref	1	Ref
RTRs	14 883	72	2.70	2.13-3.43	2.63	2.07-3.33

CI, confidence interval; GW, genital warts; HR, hazard ratio; RTR, renal transplant recipient.

^aAdjusted for age at baseline and sex.

^bAdjusted for age at baseline, sex, education, and income.

- HRs of AGWs were increased in RTRs compared with non-RTRs
 - <| year of transplanation throughout follow-up (≥</p> 10 years)
 - Independent of year of transplantation
 - In all age groups at transplantation

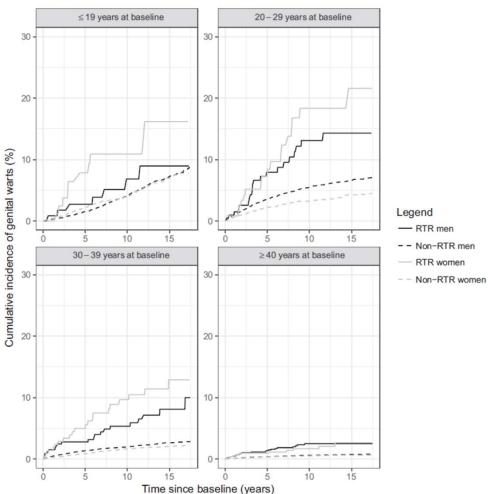
Received: 18 April 2018 | Revised: 8 July 2018 | Accepted: 25 July 2018 DOI: 10.1111/ajt.15056

ORIGINAL ARTICLE

Risk of genital warts in renal transplant recipients—A registrybased, prospective cohort study

AJT





> Acta Derm Venereol. 2021 Jul 13;101(7):adv00497. doi: 10.2340/00015555-3858.

Risk of Anogenital Warts in Renal Transplant Recipients Compared with Immunocompetent Controls: A Cross-sectional Clinical Study

Helle K Larsen ¹, Louise T Thomsen, Merete Hædersdal, Trine Thorborg Lok, Jesper Melchior Hansen, Søren Schwartz Sørensen, Susanne K Kjær

Recruitment:

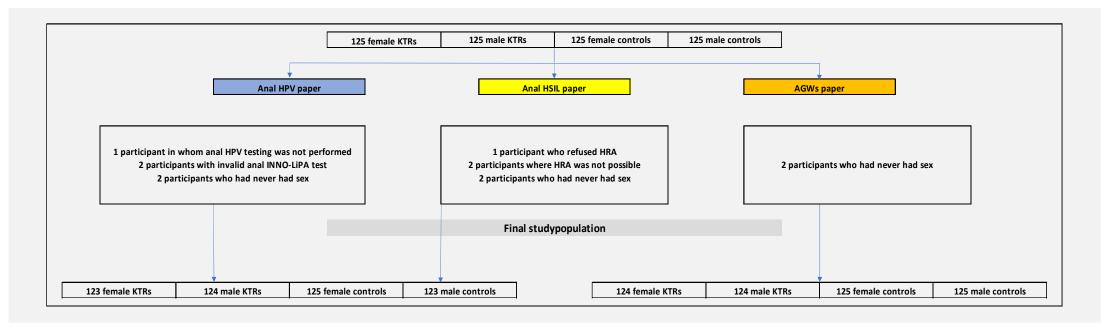
KTRs: 5 departments of Nephrology and Department of Dermato-venereology, Bispebjerg Hospital

Controls: Department of Dermato-venereology, Bispebjerg Hospital, capillary

disorders (laser), skin (pre-)cancer screening

Inclusion criterias:

- HIV uninfected
- ≥18 years of age
- no prior HPV vaccination
- no known condition requiring immunosuppressive treatment (other than the kidney transplant)
- KTRs at least 6 months posttransplant



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Definition of subclinical intra- or perianal warts: Only visible after application of acetic acid 3%



Definition of clinical AGWs: Anogenital lesions visible without application of acetic acid 3%



Histologically confirmed

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Table I. Prevalence and odds ratios (ORs) of clinical anogenital warts in renal transplant recipients and immunocompetent controls

	Men			Women					
	DTD	Controls	ORs of clinical AGV	DTDa	Control	ORs of clinical AGWs			
Anogenital site ^a	RTRs (n = 124) n (%)		Model 1 ^c OR (95% CI)	Model 2 ^d OR (95% CI)	RTRs (n=124) n (%)	Controls (n = 125) n (%)	Model 1 ^c OR (95% CI)	Model 2 ^d OR (95% CI)	
Intra-anal	4 (3.2)	8 (6.4)	0.47 (0.14-1.62)	0.31 (0.07-1.32)	11 (8.9)	9 (7.2)	1.26 (0.49-3.22)	1.04 (0.38-2.87)	
Perianal	8 (6.5)	2 (1.6)	4.42 (0.92-21.39)	4.41 (0.82-23.64)	9 (7.3)	1 (0.8)	9.85 (1.21-80.32)	7.49 (0.89-63.35)	
Vulvar	Not applica	ble			7 (5.6)	1 (0.8)	9.39 (1.11-79.76)	9.35 (1.06-82.67)	
Penile	2 (1.6)	1 (0.8)	2.04 (0.18-22.87)	1.80 (0.15-20.58)	Not applica	ble			
Any external clinical AGWsb	10 (8.1)	2 (1.6)	5.45 (1.17-25.46)	5.09 (1.03-25.04)	14 (11.3)	2 (1.6)	8.94 (1.94-41.14)	8.09 (1.69-38.82)	
Any clinical AGWs	14 (11.3)	10 (8.0)	1.45 (0.62-3.41)	1.27 (0.49-3.28)	21 (16.9)	11 (8.8)	2.10 (0.95-4.62)	1.94 (0.86-4.37)	

^aNumbers for separate sites may not sum to total, because some participants had AGWs at more than 1 site. ^bIncludes perianal, vulvar and penile clinical anogenital warts (AGWs). ^cAdjusted for age; with immunocompetent controls as the reference group. ^dAdjusted for age, smoking, lifetime number of sexual partners and (except in models for penile and vulvar warts) receptive anal sex.

- Prevalence of subclinical perianal warts: men (KTRs 9.7% vs controls 8.8%) and women (KTRs 6.5% vs controls 5.6%).
- Only a few participants had subclinical intra-anal warts (1.6% of KTRs vs 0.4% of controls)

^{95%} CI: 95% confidence interval; OR: odds ratio, RTRs: renal transplant recipients.

Received: 18 April 2018 | Revised: 8 July 2018 | Accepted: 25 July 2018 DOI: 10.1111/ajt.15056

AJT

ORIGINAL ARTICLE

Risk of genital warts in renal transplant recipients—A registrybased, prospective cohort study

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Consequence.....

KTRs/SOTRs attending regular screening for cutaneous (pre-) cancer lesions at Departments of Dermatology in Denmark are now offered inspection of the external anogenital area for warts

Project overview									
Registry-based study	Kidney transplan	t recipients	Clinical study						
Risk of genital warts		Prevalence and risk	of anogenital warts						
Risk of cervical-, vulvar-, vaginal-, penile- and anal precancerous lesions and cancer		Prevalence and risk	of anal HPV infection						
and anal precancerous lesions and cancer		Prevalence and risk	of anal HSIL						
		Prevalence of cervice infektion	al HPV infection and risk of anal co-						

> Clin Infect Dis. 2022 Nov 30;75(11):1993-1999. doi: 10.1093/cid/ciac285.

Anal Human Papillomavirus Infection in Kidney Transplant Recipients Compared With Immunocompetent Controls

Helle Kiellberg Larsen ^{1 2}, Susanne K Kjaer ^{1 3}, Merete Haedersdal ², Alexander K Kjaer ¹, Jesper Hansen Bonde ⁴, Søren Schwartz Sørensen ⁵, Louise T Thomsen ¹

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	Male KTRs vs.	male controls	Female KTRs vs. female controls				
	Prevalence,	OR ¹ (95%CI)	Prevalence	OR ¹ (95%CI)			
Anal hrHPV ²	19.4% vs. 23.6%	0.85 (0.44–1.64)	45.5% vs. 27.2%	2.87 (1.57–5.22)			

¹Adjusted for age (linear), lifetime number of sexual partners (linear), current smoking (yes/no) and receptive anal sex (ever/never) with immunocompetent controls as the reference group

²INNO-LiPA® HPV Genotyping Extra II assay (Fujirebio). HPV types 16, 18, 31, 33, 35, 39, 45, 51, 52, 56, 58, 59 and 68 were considered hrHPV types.

> Clin Infect Dis. 2022 Nov 30;75(11):1993-1999. doi: 10.1093/cid/ciac285.

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HPV-51 was the most prevalent hrHPV type in all 4 groups.

HPV-16 was found in approximately 16%–17% of the hrHPV-positive KTRs and 24%–29% of the hrHPV-positive controls.

Among hrHPV positive KTRs, 63% and 52% of men and women, respectively, were infected with hrHPV types covered by the nonavalent HPV vaccine (16/18/31/33/45/52/58).

> Transpl Infect Dis. 2023 Apr;25(2):e14019. doi: 10.1111/tid.14019. Epub 2023 Feb 7.

Prevalence of cervical human papillomavirus and the risk of anal co-infection in kidney transplant recipients: Results from a Danish clinical study

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Linea Landgrebe Ring <sup>1</sup>, Louise T Thomsen <sup>1</sup>, Merete Haedersdal <sup>2</sup>, Søren Schwartz Sørensen <sup>3</sup>, Jesper Hansen Bonde <sup>4</sup>, Trine Thorborg Lok <sup>4</sup>, Helle K Larsen <sup>2</sup>, Susanne K Kjaer <sup>1 5</sup>
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Table 3. Prevalence and odds ratio of cervical and anal hrHPV co-infection in Danish kidney transplant recipients and immunocompetent controls, 2016–2017.

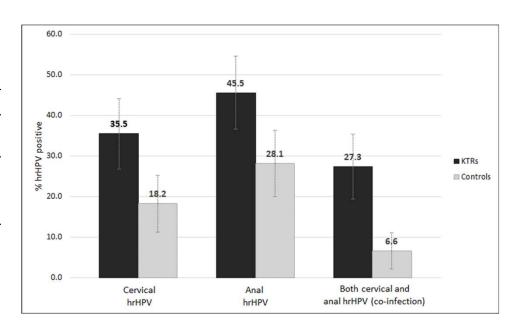
	n (%) with cervical and anal				lel 1ª	Mod	lel 2 ^b
Group	N	hr	·HPV co- nfection	OR	(95% CI)	OR	(95% CI)
Control s	121	8	(27.3)	1	(ref)	1	(ref)
KTRs	121	33	(6.6)	5.7	(2.5– 13.3)	6.3	(2.7– 15.0)

CI, confidence interval; hrHPV, high-risk human papillomavirus; OR, odds ratio; KTR, kidney transplant recipient. ^aAdjusted for age (linear).

> Transpl Infect Dis. 2023 Apr;25(2):e14019. doi: 10.1111/tid.14019. Epub 2023 Feb 7.

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^bAdjusted for age (linear), lifetime number of sexual partners (linear), current smoking (yes/no) and history of receptive anal sex (yes/no).

> Transpl Infect Dis. 2023 Apr;25(2):e14019. doi: 10.1111/tid.14019. Epub 2023 Feb 7.

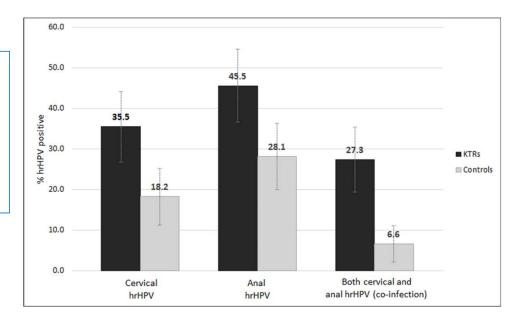
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30.3% (10/33) had the exact same HPV types detected at both sites;

33.3% (11/33) had both concordant and discordant HPV types on the cervix and anus; and

36.4% (12/33) did not have any of the same HPV types detected at the two sites



Project overview								
Registry-based study	Kidney transplan	t recipients	Clinical study					
Risk of genital warts		Prevalence and risk	of anogenital warts					
Risk of cervical-, vulvar-, vaginal-, penile- and anal precancerous lesions and cancer		Prevalence and risk	of anal HPV infection					
and anal precamed out resions and cancer		Prevalence and risk	of anal HSIL					
		Prevalence of cervic	al HPV infection and risk of anal co-					

infektion

Human papillomavirus-related anogenital premalignancies and cancer in renal transplant recipients: A Danish nationwide, registry-based cohort study

Kristian Reinholdt 1 , Louise T Thomsen 1 , Christian Dehlendorff 2 , Helle K Larsen 1 3 , Søren S Sørensen 4 , Merete Haedersdal 3 , Susanne K Kjaer 1 5

Table 2. Hazard ratio (HR) of HPV-related anogenital intraepithelial neoplasia grade 2/3 or cancer in Danish female renal transplant recipients (1990–2015) vs. a nontransplanted comparison cohort (non-RTRs)

	Intraepithelia	Intraepithelial neoplasia grades 2/3							Cancer					
Anogenital site	Person-years	Events	HR¹	(95% CI)	HR ²	(95% CI)	Person-years	Events	s HR¹	(95% CI)	HR ²	(95% CI)		
Cervical														
Non-RTRs	910,648	1,542	1	(ref)	1	(ref)	964,349	211	1	(ref)	1	(ref)		
RTRs	14,017	59	2.1	(1.7-2.8)	2.1	(1.7-2.8)	15,055	9	2.8	(1.4-5.4)	2.7	(1.4-5.3)		
Vaginal														
Non-RTRs	964,071	14	1	(ref)	1	(ref)	964,349	15	1	(ref)	1	(ref)		
RTRs	15,009	7	32.7	(13.1-81.6)	35.0	(13.9-87.7)	15,055	< 5	11.1	(2.5-49.4)	10.7	(2.4-47.9)		
Vulvar														
Non-RTRs	963,032	92	1	(ref)	1	(ref)	964,349	20	1	(ref)	1	(ref)		
RTRs	14,936	24	16.8	(10.7-26.4)	16.4	(10.4-25.8)	15,055	8	31.7	(13.7-73.4)	31.0	(13.3-72.0)		
Anal														
Non-RTRs	964,019	26	1	(ref)	1	(ref)	964,349	38	1	(ref)	1	(ref)		
RTRs	14,991	19	48.8	(26.9-88.8)	51.1	(28.0-93.1)	15,055	5	10.0	(3.9-25.4)	10.4	(4.1-26.5)		

Human papillomavirus-related anogenital premalignancies and cancer in renal transplant recipients: A Danish nationwide, registry-based cohort study

Kristian Reinholdt ¹, Louise T Thomsen ¹, Christian Dehlendorff ², Helle K Larsen ¹ ³, Søren S Sørensen ⁴, Merete Haedersdal ³, Susanne K Kjaer ¹ ⁵

Table 3. Hazard ratio (HR) of HPV-related anogenital intraepithelial neoplasia Grade 2/3 or cancer in Danish male renal transplant recipients (1990–2015) vs. a nontransplanted comparison cohort (non-RTRs)

	Intraepithelial	neoplas	ia grad	les 2/3			Cancer					
Anogenital site	Person-years	Events	HR1	(95% CI)	HR ²	(95% CI)	Person-years	Events	HR1	(95% CI)	HR ²	(95% CI)
Penile												
Non-RTRs	1,608,284	36	1	(ref)	1	(ref)	1,608,628	52	1		1	
RTRs	25,469	11	22.7	(11.5-45.0)	21.9	(11.1-43.5)	25,531	6	9.5	(4.1-22.3)	9.6	(4.1-22.4)
Anal												
Non-RTRs	1,608,542	14	1	(ref)	1	(ref)	1,608,628	31	1		1	
RTRs	25,491	9	38.9	(16.7-90.6)	39.0	(16.7-91.1)	25,531	0	NA		NA	
Penile/anal comb	bined											
Non-RTRs	1,608,198	50	1	(ref)	1	(ref)	1,608,628	83	1		1	
RTRs	25,429	20	27.7	(16.4-46.9)	26.9	(15.9-45.5)	25,531	6	5.6	(2.4-12.9)	5.6	(2.4-12.9)

¹Adjusted for age at baseline.

²Adjusted for age at baseline, income and educational level.

> Clin Infect Dis. 2021 Jul 1;73(1):21-29. doi: 10.1093/cid/ciaa781.

Risk of Anal High-grade Squamous Intraepithelial Lesions Among Renal Transplant Recipients Compared With Immunocompetent Controls

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Helle K Larsen <sup>1</sup> <sup>2</sup>, Merete Hædersdal <sup>2</sup>, Louise T Thomsen <sup>1</sup>, Rasmus Hertzum-Larsen <sup>1</sup>, Trine Thorborg Lok <sup>3</sup>, Jesper Bonde <sup>3</sup>, Søren S Sørensen <sup>4</sup>, Jesper Melchior Hansen <sup>5</sup>, Joel M Palefsky <sup>6</sup>, Susanne K Kjær <sup>1</sup> <sup>7</sup>
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Odds ratios of anal HSIL in kidney transplant recipients compared with controls

	total stu	dy population	Anal high-risk HPV positive participants				
	n(%)	_	n(%)				
	anal HSIL	OR* (95% CI)	anal HSIL	OR* (95% CI)			
Men							
controls	(8.0)	1 (ref)	(3.4)	1 (ref)			
KTRs	(6.5)	11.21 (1.46–291.17)	(33.3)	14.84 (1.65–438.24)			
Women							
Controls	(4.0)	1 (ref)	(14.7)	1 (ref)			
KTRs	(15.4)	6.41 (2.14–24.10)	(33.9)	4.58 (1.36–18.85)			

^{*}Adjusted for age, smoking, lifetime number of sexual partners and receptive anal sex.

CI, confidence interval, HSIL, high-grade intraepithelial lesion; OR, odds ratio; KTR, kidney transplant recipient

Anal HSIL was exclusively found in anal hrHPV positive KTRs and controls

> Clin Infect Dis. 2021 Jul 1;73(1):21-29. doi: 10.1093/cid/ciaa781.

Risk of Anal High-grade Squamous Intraepithelial Lesions Among Renal Transplant Recipients Compared With Immunocompetent Controls

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HPV 16 and anal HSIL:

Prevalence of aHSIL in individuals with HPV16 infection:

53.8% (7/13) in KTRs versus 29.4% (5/17) in controls (p=0.26).

For other hrHPV types than HPV16, the corresponding prevalences of aHSIL were

29.9% (20/67) in KTRs versus 2.2% (1/46) in controls (p<0.001).

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RISK FACTORS INTRA-/PERIANAL WARTS AND ANAL HSIL

RTRs	Intra-/perianal warts, OR ¹ (95%CI)	Anal HSIL, OR¹ (95% CI)
Receptive anal sex	2.60 (1.11-6.07)	6.23 (2.23-19.08)
Number of lifetime sexual partners ≥ 10	1.90 (0.82-4.40)	3.04 (0.84-11.88)
Current smoking	2.43 (1.10-5.39)	0.89 (0.22-2.89)
History of AGWs	1.04 (0.43-2.48)	4.21 (1.53-11.48)
Anal hrHPV infection		

^{&#}x27;Adjusted for age, sex, smoking, number of lifetime sexual partners and receptive anal sex.

²Fisher's exact test: p<0.0001, OR not computable as there were no cases of anal HSIL in anal hrHPV negative RTRs

CASE I FROM THE KTR HPV/DYPLASIA PROJECT

- 36-year old female KTR, Kidney transplatation maj 2010
 Cause: Polycystic kidney disease
- Immunosuppressive treatment:
 - Prednisolon
 - Tacrolimus (prograf)
 - Mycophenolatmofetil

Clinical examnation 2016:

Verrucous lesion at the introitus

Histologically: Vulva HSIL-VIN3

Cervix: persistent infection with HPV 16



CASE I : ANAL FINDINGS

Anal cytology: ASC-H

Anal HPV testing: HPV 16

HRA: 9 o'clock mosaicism, 4 o'clock leucoplakia with punctuaion, erosive

Histology: Anal HSIL-AIN3 x 2





CASE I CONTINUED

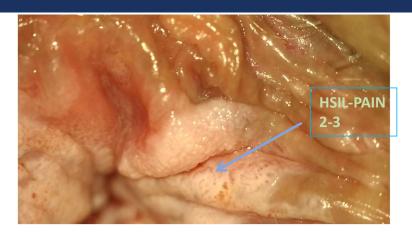
- Ongoing anal HSIL and
- From 2020 perianal HSIL
- Treatments
 - 5-FU, Imiquimod 5%, Trichloroacetic acid
 - Electrocautery x 5
 - From dec 2022 referred to department of Gastrointestinal surgery

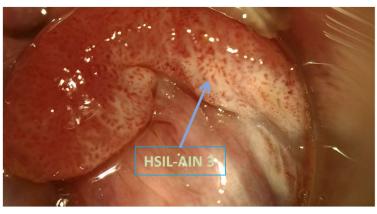
September 2023 Perianal SCC

CASE II FROM THE KTR HPV/DYPLASIA PROJECT

71-YEAR OLD FEMALE KTR, ANAL CYTOLOGY: ASC-H, ANAL HPV: HPV 16



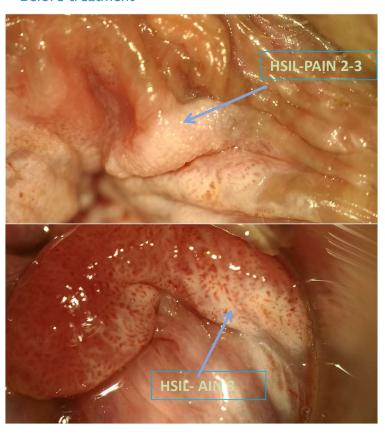






CASE 2 BEFORE AND AFTER TREATMENT

Before treatment



After treatment: electro cautery x many + surgical removal of polyp





CASE III 47-YEAR OLD KIDNEY TRANSPLANT RECIPIENT, REFERRED AFTER PERIANAL SCC X 2

Systemic Lupus Erythematosus since 2000 Kidney transplant: 2014

Perianal SCC x 2; August 2018 and January 2020.



CASE III CONTINUED



Histology:

Perianal:

12.07.2018 **HSIL** at 5-6 o'clock, hrHPV type 18 og 33

14.08.2018 SCC at 5-6 o'clock, free resectionborder, PET CT uden tegn til spredning. No indication for X-ray therapy

05.12.2018 **HSIL** at 3-5 o'clock, excision with el-diathermy

23.08-2019, **HSIL** at 4, 4-5, 6-7 and 7 o'clock

30-01-2020 SCC recurrence perianal at 5 o'clock. X-ray treatment 27/30 54 gy for T1 c ani 03.06.2020 **HSIL** at 4 and 8 o'cloco.

26.11.2021 og 10.03.2022 **HSIL** at 4 o´clock

16.06.2022 **HSIL** at 6 0'clock

03.10.2022 **HSIL** 5 and 7 o'clock

26.01.2023 **HSIL** at 5 o'clock

24.08.2023: **HSIL** at 7 o'clock

02.01.2024 **HSIL** at 5 o'clock, at 6-7 o'clock changes corresponding to HSV, confirmed on immunohistochemical staining

04.06.2024 **HSIL** at 3, 5-6 and 6 o'clock.

Analcanal:

19.12.2019 **HSIL** at 11 o'clock and LSIL at 8 o'clock.

29.09.2022 HSIL at 6 o'clock and probably HSIL at 8-9 o'clock

Vulva:

26.01.2023 Introitus **HSIL**

20.04.2023 HSIL lateral for lleft labia major

Cervix

2013 Conisatio: CIN3.

2020 hrHPV type 16 og 33, normal cytology (februar 2020)



Treatments perianal/anal HSIL:

Aldara 5% x 3 weekly for 8 weeks only slight effect

5-FU only one week due to sideeffects.

Trichloroacetic acid 90% x many

Excision with diathermy dept of Surgery 16.06.2022 and sept 2022

Electrocautery x 4

Cryotherapy 2 x 20 sec 04.07.2024 og 3 x 10 sec $\,$ 11.07.2024 followed by topical 5-FU x 5 weekly for $\,$ 6+4 weeks with good result

CASE III CONTINUED BEFORE AND AFTER CRYO FOLLOWED BY TOPICAL 5-FU





CASE III CONTINUED ON DIALYSIS, ONLY STEROID TREATMENT CURRENTLY

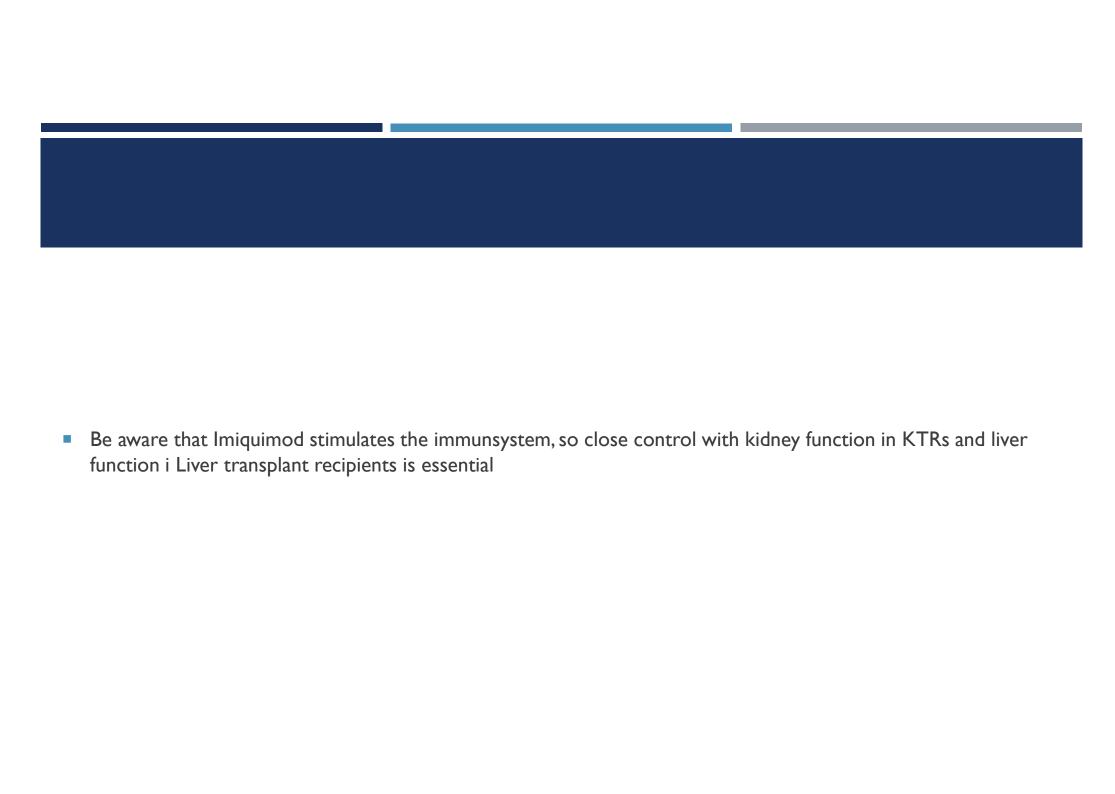






Conclusions and Perspectives

- In a Danish population of kidney transplant recipients
 - Increased prevalence and risk of anogenital warts compared to healthy controls were found
 - At Dermatological visits for skin cancer screening a full skin examination should include inspection of the anogenital area for anogenital warts



Conclusions and Perspectives

- In a Danish population of kidney transplant recipients compared to healthy controls:
 - Increased prevalence and risk of anogenital warts
 - At Dermatological visits for skin cancer screening a full skin examination shall include inspection of the anogenital area for anogenital warts
 - Higher prevalence and risk of anal hrHPV in female KTRs
 - For both male and female KTRs with anal hrHPV similar prevalence of anal HSIL (33%)
 - >50% of KTRs with anal HPV16 had anal HSIL and aprox. 30% of KTRs with other anal hrHPV than HPV 16 had anal HSIL
 - Anal hrHPV seems reasonable as a triage marker for HRA referral in KTRs

SPECIAL REPORT



International Anal Neoplasia Society's consensus guidelines for anal cancer screening

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Elizabeth A. Stier 10 | Megan A. Clarke 20 | Ashish A. Deshmukh 3,40 |
Nicolas Wentzensen 20 | Yuxin Liu 50 | I. Mary Poynten 60 |
Eugenio Nelson Cavallari 7 | Valeria Fink 8 | Luis F. Barroso 9 |
Gary M. Clifford 100 | Tamzin Cuming 11 | Stephen E. Goldstone 12 |
Richard J. Hillman 6,13 | Isabela Rosa-Cunha 14 | Luciana La Rosa 15,16 |
Joel M. Palefsky 17 | Rosalyn Plotzker 18 | Jennifer M. Roberts 190 | Naomi Jay 17
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TABLE 1 Populations for screening.

Population—Risk category	When	Anal cancer incidence ^{2,5} per 100,000 person-years
Risk Category A (incidence ≥ 10-fold compared to the general population	n)	
MSM and TW with HIV	Age 35	>70/100,000 age 30-44 >100/100,000 age 45+
Women with HIV	Age 45	>25/100,00 age 45+
MSW with HIV	Age 45	>40/100,000 age 45+
MSM and TW not with HIV	Age 45	>18/100,000 age 45-59 >34/100,000 age 60+
History of vulvar HSIL or cancer	Within 1 year of diagnosis	>40/100,000
Solid organ transplant recipient	10 years post-transplant	>25/100,000
Risk Category B (incidence up to 10-fold higher compared to the general	al population)	
Cervical/vaginal cancer	Shared decision age 45 ^a	9/100,000
Cervical/vaginal HSIL	Shared decision age 45 ^a	8/100,000
Perianal warts (male or female)	Shared decision age 45 ^a	Unknown
Persistent cervical HPV 16 (>1 year)	Shared decision age 45 ^a	Unknown
Other immunosuppression (e.g., Rheumatoid arthritis, Lupus, Crohn's, Ulcerative colitis, on systemic steroid therapy)	Shared decision age 45 ^a	6/100,000
Incidence among the general population: 1.7 per 100,000 ⁸		



International Anal Neoplasia Society's consensus guidelines for anal cancer screening

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Elizabeth A. Stier<sup>1</sup> | Megan A. Clarke<sup>2</sup> | Ashish A. Deshmukh<sup>3,4</sup> | Nicolas Wentzensen<sup>2</sup> | Yuxin Liu<sup>5</sup> | I. Mary Poynten<sup>6</sup> | Eugenio Nelson Cavallari<sup>7</sup> | Valeria Fink<sup>8</sup> | Luis F. Barroso<sup>9</sup> | Gary M. Clifford<sup>10</sup> | Tamzin Cuming<sup>11</sup> | Stephen E. Goldstone<sup>12</sup> | Richard J. Hillman<sup>6,13</sup> | Isabela Rosa-Cunha<sup>14</sup> | Luciana La Rosa<sup>15,16</sup> | Joel M. Palefsky<sup>17</sup> | Rosalyn Plotzker<sup>18</sup> | Jennifer M. Roberts<sup>19</sup> | Naomi Jay<sup>17</sup>
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"In settings with limited HRA capacity, it is acceptable to only refer individuals testing positive for HPVI6 to immediate HRA, with repeat testing in I2 months recommended for individuals testing positive for other hrHPV types. Repeat screening in 24 months is recommended for individuals testing hrHPV negative".

CORE STUDY GROUP

Dept. of Dermatology, Bispebjerg Hospital

Study design, funding, coordination, patient recruitment, data collection, publication.

Team members:

Responsibilities:

Helle Kiellberg Larsen (MD, PhD, ass. Professor) Merete Hædersdal (professor, MD, DMSc.) Benedikte Skov (research nurse)

Expertise in dermatology and skin cancer in SOTRs.





Danish Cancer Society Research Center

Team members:

Susanne Krüger Kjær (professor, MD, DMSc.)



Helle Kiellberg Larsen (MD, PhD, ass professor), employed parttime at time of study Linea Landgrebe Ring, PhD student

Christian Dehlendorff, Head of statistics (PhD)

Responsibilities:

Expertise in HPV, cervical cancer, clinical epidemiology and biostatistics.

Study design, funding, data management, registry linkage, statistical analysis, publication.





COLLABORATORS

Expertise in organ transplantation

Dept. of Nephrology, Rigshospitalet Søren Schwartz Sørensen (professor, MD, DMSc.)

Dept. of Nephrology, Herlev Hospital

lesper Melchior Hansen, consultant, MD, DMSc

External assessor on anal HSIL paper

UCSF Anal dysplasia clinic, San Francisco

Professor Joel Palefsky

HPV testing, Cytology, Histology

Dept. of Pathology, Hvidovre Hospital

Trine Thorborg Lok, (consultant at time of study)

Jesper Bonde, senior researcher

Helle Pedersen, biomedical laboratory scientist

