

Appendix. Two-year Study History of 33 Women Diagnosed With CIN-3 During ALTS Who Tested Negative for Carcinogenic HPV at Baseline

	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 1 Clinical center=AL Study arm=IC Age at enrollment=21	-1	Referral	LSIL	LSIL									
	0	0 (with colpo)			P1	LSIL	LSIL	LSIL/CIN-1	Neg	Low grade	Neg	Neg	
	5	6		Neg	P1	Neg	Neg				Neg		
	11	12			Atypical	Neg					Pos		
	18	18			P1	LSIL	ASC-US				Pos	<u>16-40-54-56-67-70-84</u>	
	24	24 (with colpo)				ASC-US	ASC-US	CIN-2	CIN-3	Low grade	Pos	06- <u>16-18-66-73</u>	
	25	Excision						LSIL/CIN-1	LSIL/CIN-1				
Subject 2 Clinical center=AL Study arm=CM Age At enrollment=22	-2	Referral	ASC-US	ASC-US									
	0	0			Neg	Neg	ASC-US				Neg	Neg	18-33
	9	6		Neg	Neg	Neg	Neg				Neg	Neg	
	12	12*											
	17	18			Neg	Neg					Pos	62	
	24	24 (with colpo)			Neg	ASC-US	HSIL	CIN-2	CIN-3	Low grade	Neg	33	
	26	Excision						ASC	ASC				
Subject 3 Clinical center=AL Study arm=HPV Age at enrollment=24	-2	Referral	ASC-US	ASC-US									
	0	0		Neg	Neg	Neg	Neg				Neg		<u>58-59-70-Mix</u>
	6	6			Neg	LSIL	LSIL				Pos	<u>52-58</u>	
	12	12			Neg	UNSAT					Pos	Neg	
	13	12			ND	Neg					ND	<u>18-35-42-59-89</u>	
	18	18			Neg	LSIL					Pos	18-35	
	25	24 (with colpo)			Neg	LSIL	LSIL	LSIL/CIN-1	LSIL/CIN-1	Low grade	Pos	35-71	
	27	Excision						CIN3	CIN3				

Online appendix to Castle PE, Cox T, Jeronimo J, Solomon D, Wheeler CM, Gravitt PE et al. An Analysis of High-Risk Human Papillomavirus DNA Negative Cervical Precancers in the ASC-US-LSIL Triage Study (ALTS). Obstet Gynecol 2008;111:xxx-xx.

	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 4 Clinical center=AL Study arm=HPV Age At enrollment=22	-3	Referral	LSIL	LSIL									
	0	0			P2	LSIL	LSIL				Neg	26	
	6	6*											
	12	12*											
	18	18*											
	25	24 (with colpo)			P1	HSIL	HSIL	CIN-2	CIN-2	High grade	Neg	26	
	27	Excision						CIN-3	CIN-3				
Subject 5 Clinical center=AL Study arm=IC Age At enrollment=19	-1	Referral	ASC-US	ASC-US									
	0	0 (with colpo)			Atypical	Neg	ASC-US	Neg	Neg	Low grade	Neg	Neg	<u>51</u>
	5	6			Neg	Neg	LSIL		Neg		Pos		
	12	12			P1	LSIL					Pos	<u>16-51-59</u>	
	18	18*											
	23	24 (with colpo)			P1	LSIL	LSIL	ASC	Neg	Low grade	Pos	<u>16-51-59</u>	
	25	colpo						CIN-2	CIN-3	Low grade			
Subject 6 Clinical center=AL Study arm=CM Age at enrollment=18	27	Excision						CIN-2	CIN-3				
	-2	Referral	LSIL	LSIL									
	0	0			Neg	Neg	Neg				Neg	Neg	
	6	6		Neg	Neg	LSIL	ASC-US				Pos		
	12	12			P1	LSIL					Pos	06-42-52	
	18	18			P1	Neg					Pos		
	24	24 (with colpo)			P1	Neg	ASC-US	ASC	CIN-3	Low grade	Pos	Neg	
	29	Excision						CIN-2	CIN-2				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 7 Clinical center=AL Study arm=HPV Age at enrollment=18	-6	Referral	ASC-US	LSIL									
	0	0			Neg	Neg	ASC-US				Neg	55-81	55-81
	6	6*											
	12	12*											
	18	18*											
	25	24 (with colpo)			Neg	HSIL	HSIL	LSIL/CIN-1	CIN-2	Low grade	Pos	35	
	26	Excision						CIN-3	CIN-2				
Subject 8 Clinical center=AL Study arm=HPV Age at enrollment=28	-5	Referral	ASC-US	ASC-US									
	0	0			Neg	Neg	Neg				Neg	Neg	<u>68</u>
	6	6			Neg	Neg					ND	<u>16-66-68</u>	
	12	12			Neg	ASC-US					Pos	<u>16-66</u>	
	18	18			Neg	ASC-US	ASC-US				ND	<u>16</u>	
	25	24 (with colpo)			Neg	ASC-US	ASC-US	LSIL/CIN-1	LSIL/CIN-1	Low grade	ND	<u>16</u>	
	28	Excision						CIN-2	CIN-3				
Subject 9 Clinical center=AL Study arm=IC Age at enrollment=24	-2	Referral	ASC-US	Neg									
	0	0 (with colpo)			Atypical	Neg	Neg	ND		normal	Neg	42	42-45-62
	6	6			Neg	LSIL					Pos	58-83	
	12	12			Atypical	LSIL	LSIL				Pos	<u>35</u>	
	18	18			Neg	ASC-US	LSIL				Pos	<u>35</u>	
	24	24 (with colpo)			Neg	LSIL	LSIL	LSIL/CIN-1	LSIL/CIN-1	Low grade	Pos	Neg	
	26	Excision						CIN-3	CIN-3				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 10 Clinical center=AL Study arm=IC Age at enrollment=32	-1	Referral	ASC-US	ASC-US									
	0	0 (with colpo)			P1	Neg	Neg	LSIL/CIN-1	CIN-3	Normal	Neg	Neg	58-Mix
Subject 11 Clinical center=OK Study arm=IC Age at enrollment=24	-3	Referral	ASC-US	LSIL									
	0	0 (with colpo)			Neg	Neg	Neg	ND		Normal	Neg	Neg	
	6	6			Neg	Neg	Neg				ND	<u>52</u>	
	14	12			Neg	ASC-US					Pos	<u>33-52</u>	
	20	18			Neg	ASC-US					Pos	<u>33-52</u>	
	25	24 (with colpo)			Neg	HSIL	HSIL	CIN-2	CIN-2	Low grade	Pos	<u>33-52</u>	
	28	Excision						CIN-3	CIN-3				
Subject 12 Clinical center=OK Study arm=IC Age at enrollment=38	-2	Referral	LSIL	ASC-US									
	0	0 (with colpo)			Neg	Neg	ASC-US	LSIL/CIN-1	Neg	Low grade	Neg	Neg	
	6	6			P1	Neg					Neg		
	12	12			Neg	Neg	ASC-US				Pos	<u>67</u>	
	18	18			P1	ASC-US	LSIL				Pos	<u>67</u>	
	25	24 (with colpo)			P1	Neg	Neg	CIN-2	CIN-3	Low grade	Neg		
	28	Excision						LSIL/CIN-1	Neg				
Subject 13 Clinical center=OK Study arm=IC Age at enrollment=21	-2	Referral	LSIL										
	0	0			Neg	Neg	Neg				Neg	Neg	
	1	Colpo						CIN-3	ASC	Low grade			
	2	Excision						Neg	Neg				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 14 Clinical center=PA Study arm=IC Age at enrollment=21	-1	Referral	LSIL										
	0	0 (with colpo)			Atypical	Neg	Neg	Neg	Neg	Metaplasia	Neg	Neg	
	6	6*											
	12	12			Neg	Neg					Pos	<u>16-31-33-52-53-54-59-62-89</u>	
	18	18*											
	24	24 (with colpo)			Neg	ASC-US	Neg	CIN-2	CIN3	Low grade	Pos	<u>16-31-33-40-52-54-59</u>	
	28	Excision						Neg	Neg				
Subject 15 Clinical center=PA Study arm=IC Age at enrollment=27	-3	Referral	LSIL	HSIL									
	0	0 (with colpo)			P1	LSIL	LSIL	LSIL/CIN-1	CIN-2	Low grade	ND	Neg	
	6	6*											
	12	12*											
	18	18*											
	27	24 (with colpo)			P1	HSIL	HSIL	CIN-2	CIN-3	Low grade	Pos	16-54	
	29	Excision						LSIL/CIN-1	Neg				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 16 Clinical center=PA Study arm=CM Age at enrollment=19	-4	Referral	ASC-US	ASC-US									
	0	0			P1	Neg	Neg				ND	84	84
	7	6			Neg	ASC-US					Pos	<u>51-55-81</u>	
	12	12			P1	LSIL	LSIL				Pos	40- <u>51-55-81</u>	
	19	18			Atypical	LSIL	HSIL				Pos	<u>16-55-61-81</u>	
	24	24 (with colpo)			P1	LSIL	LSIL	LSIL/CIN-1	LSIL/CIN-1	Low grade	ND	<u>16</u>	
	26	Excision						CIN-2	CIN-3				
Subject 17 Clinical center=PA Study arm=IC Age at enrollment=36	-2	Referral	LSIL	HSIL									
	0	0 (with colpo)			Neg	HSIL	Neg	CIN-3	CIN-3	High grade	Neg	Neg	
	1	Excision						CIN-3	CIN-3				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 18 Clinical center=PA Study arm=CM Age at enrollment=47	-2	Referral	ASC-US	ASC-US									
	0	0			Neg	LSIL	ASC-US				Neg	Neg	84
	6	6		ASC-US	Neg	LSIL	LSIL				ND	Neg	
	12	12			Neg	LSIL	LSIL				Pos	67-70-84-89	
	18	18			Neg	LSIL					Pos	16-70-89	
	24	24 (with colpo)			Neg	ASC-US	HSIL	CIN-2	CIN-3	Low grade	Pos	16-52-70-84-89	
	27	Excision						CIN-2	CIN-2				
	31	Colpo						CIN-2	LSIL/CIN-1	Low grade			
	34	Excision						CIN-2	CIN-2				
Subject 19 Clinical center=PA Study arm=IC Age at enrollment=34	-1	Referral	ASC-US	Neg									
	0	0 (with colpo)			Neg	Neg	Neg	ND		Normal	Neg	Neg	Neg
	4	6			Neg	Neg					Neg	Neg	
	13	12			Neg	LSIL					Pos	31	
	18	18			Atypical	HSIL	HSIL				Pos	31	
	20	Colpo						CIN-2	CIN-2	High grade			
	21	Excision						CIN-2	CIN-3				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 20 Clinical center=PA Study arm=CM Age at enrollment=51	-2	Referral	ASC-US	ASC-US									
	0	0			P1	ASC-US	ASC-US				Neg	Neg	Neg
	5	6			P1	LSIL	ASC-US				Neg	Neg	
	11	12			Neg	ASC-US	ASC-US				Neg	Neg	
	17	18			Neg	Neg	Neg				Neg	Neg	
	23	24 (with colpo)			Neg	ASC-US	ASC-US	CIN-2	CIN-3	Metaplasia	Neg	Neg	
	25	Excision						Neg	Neg				
Subject 21 Clinical center=PA Study arm=IC Age at enrollment=70	-2	Referral	ASC-US	Neg									
	0	0 (with colpo)			Neg	ASC-US	ASC-US	Neg	CIN-3	Normal	Neg	Neg	Neg
Subject 22 Clinical center=PA Study arm=IC Age at enrollment=38	-2	Referral	ASC-US	ASC-US									
	0	0			Neg	LSIL	ASC-US				Neg	Neg	Neg
	0	Colpo						ND		Normal			
	6	6		Neg	Neg	Neg	Neg				ND	Neg	
	13	12			Neg	ASC-US	ASC-US				ND	Neg	
	18	18*											
	24	24 (with colpo)			P1	LSIL	ASC-US	LSIL/CIN-1	CIN-3	Low grade	Neg	Neg	
	26	Excision						LSIL/CIN-1	LSIL/CIN-1				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 23 Clinical center=PA Study arm=IC Age at enrollment=27	-2	Referral	ASC-US	ASC-US									
	0	0 (with colpo)			P1	Neg	Neg	Neg	CIN3	Metaplasia	Neg	81	55-81
Subject 24 Clinical center=PA Study arm=IC Age at enrollment=25	-4	Referral	ASC-US	Neg									
	0	0 (with colpo)			Neg	ASC-US	Neg	Neg	Neg	Metaplasia	Neg	Neg	Neg
	5	6			Neg	Neg					ND	51-89	
	12	12			Neg	Neg					Pos	<u>18</u>	
	19	18			Neg	ASC-US	ASC-US				Pos	<u>18</u>	
	26	24 (with colpo)			P1	Neg	Neg	LSIL/CIN-1	LSIL/CIN-1	Metaplasia	ND	16-55	
	29	Excision						LSIL/CIN-1	CIN3				
Subject 25 Clinical center=WA Study arm=IC Age at enrollment=35	-1	Referral	ASC-US	ASC-US									
	0	0			Atypical	Neg	Neg				Neg	54	16-45-54
	0	Colpo						Neg	Neg	Low grade			
	6	6*											
	12	12*											
	19	18			P1	Neg					Neg	54	
	24	24 (with colpo)			Neg	ASC-US	HSIL	ASC	Neg	Low grade	Neg	89	
	27	Colpo						CIN2	CIN3	High grade			

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 26 Clinical center=WA Study arm=HPV Age at enrollment=25	-1	Referral	ASC-US	ASC-US									
	0	0			Neg	ASC-US	ASC-US				Neg	Neg	<u>39-51-53</u>
	6	6		Neg	ND	Neg	Neg				Pos	<u>39-51-54</u>	
	14	12			P1	ASC-US					Neg	<u>54-84</u>	
	18	18*											
	24	24 (with colpo)			ND	ASC-US	ASC-US	Neg	CIN3		Neg	06- <u>54-84</u>	
Subject 27 Clinical center=WA Study arm=IC Age at enrollment=20	-3	Referral	ASC-US	ASC-US									
	0	0 (with colpo)			Neg	ASC-US	ASC-US	CIN-3	Neg	Low grade	Neg	Neg	66
Subject 28 Clinical center=WA Study arm=IC Age at enrollment=23	-3	Referral	ASC-US	ASC-US									
	0	0 (with colpo)			Neg	Neg	ASC-US	CIN-2	CIN-3	Low grade	Neg	81	81
Subject 29 Clinical center=WA Study arm=IC Age At enrollment=21	-2	Referral	ASC-US	ASC-US									
	0	0 (with colpo)			Neg	HSIL	Neg	HSIL	LSIL/CIN-1	High grade	Neg	42-67	42-67
	1	Excision						CIN-3	CIN-3				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 30 Clinical center=WA Study arm=IC Age at enrollment=19	-5	Referral	ASC-US										
	0	0 (with colpo)			Atypical	Neg	Neg	ASC	Neg	Metaplasia	Neg	Neg	Neg
	7	6			P1	ASC-US					Neg	Neg	
	11	12			P1	ASC-US					Neg	Neg	
	18	18			P1	Neg	ASC-US				Neg	Neg	
	25	24 (with colpo)			P1	HSIL	HSIL	Neg	Neg	Metaplasia	Pos	56	
	28	Colpo						ASC	Neg	Metaplasia			
	30	Colpo						CIN3	CIN2	Low grade			
Subject 31 Clinical center=WA Study arm=CM Age at enrollment=23	-3	Referral	ASC-US	ASC-US									
	0	0			Neg	ASC-US	ASC-US				ND	Neg	84
	4	6			Neg	Neg					Neg	Neg	
	12	12			Atypical	Neg					Neg	Neg	
	18	18			P1	ASC-US					Neg	Neg	
	24	24 (with colpo)			Neg	Neg	Neg	CIN-2	Neg	Low grade	ND		
	25	Excision						LSIL/CIN-1	CIN-3				
Subject 32 Clinical center=WA Study arm=CM Age at enrollment=26	-1	Referral	ASC-US	ASC-US									
	0	0			Neg	Neg	Neg				Neg	67	61-67
	7	6			P1	HSIL	HSIL				Pos	61-67	
	9	Colpo						CIN-3	CIN-3	Low grade			
	10	Excision						Neg	Neg				

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	Time (months)	Visit Type	Pap (CC)	Pap (QC)	Cervicography Impression	LBC (CC)	LBC (QC)	Histology (CC)	Histology (QC)	Colposcopy Impression	HC2	LBA Results	LA Results
Subject 33 Clinical center=WA Study arm=CM Age at enrollment=22	-1	Referral	ASC-US	ASC-US									
	0	0			Neg	LSIL	LSIL				ND	42	42
	6	6			Neg	ASC-US	ASC-US				Pos	06- <u>16-51-66</u>	
	12	12			P1	HSIL	HSIL				ND	<u>16-51-66</u>	
	13	Colpo						CIN-2	CIN-2	High grade			
	16	Excision						CIN-3	CIN-3				

CIN, cervical intraepithelial neoplasia; ALTS, ASCUS-LSIL Triage Study; HPV, human papillomavirus; Pap, Pap smear; CC, clinical center pathology; QC, quality control pathology; LBC, liquid-based cytology; HC2, hybrid capture 2; LBA, line blot assay; LA, linear assay; AL, Alabama; IC, immediate colposcopy; LSIL, low-grade squamous intraepithelial lesion; Colpo, colposcopy; P1, low-grade colposcopic impression; Neg, negative; Pos, positive; ASC-US, atypical squamous cells of undetermined significance; CM, conservative management; HSIL, high-grade squamous intraepithelial lesion; ASC, atypical squamous cell; UNSAT, unsatisfactory; ND, not done; P2, high-grade colposcopic impression; HSIL, high-grade squamous intraepithelial lesion; OK, Oklahoma; PA, Pennsylvania; Metaplasia, atypical metaplasia.

Bolded, red type highlights CIN-3 histologic diagnoses. Underlined carcinogenic HPV types indicate detection at consecutive visits.

*Missed visit.

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