

# NIH Public Access

Author Manuscript

Sex Transm Dis. Author manuscript; available in PMC 2009 December 1.

Published in final edited form as:

Sex Transm Dis. 2008 December ; 35(12): 1011-1014. doi:10.1097/OLQ.0b013e318182c91f.

## Awareness of Kaposi's Sarcoma-associated Herpesvirus among Men who Have Sex with Men

Abigail M. Phillips<sup>1</sup>, Alison Graves Jones, M.P.H.<sup>1</sup>, Dennis H. Osmond, Ph.D.<sup>1</sup>, Lance M. Pollack, Ph.D.<sup>1</sup>, Joseph A. Catania, Ph.D.<sup>1,\*</sup>, and Jeffrey N. Martin, M.D., M.P.H.<sup>1</sup>

1 University of California, San Francisco

### Abstract

**Background**—Despite burgeoning scientific knowledge about Kaposi's sarcoma-associated herpesvirus (KSHV), the etiologic agent of Kaposi's sarcoma (KS), little is known about awareness of this virus in the general community. This is particularly the case for men who have sex with men (MSM), the group at greatest risk for infection.

**Methods**—The California Health Interview Survey was a random digit-dial survey of over 50,000 households. Men age 18–64 years who self-identified as gay or bisexual were subsequently recontacted for a follow-up study of HIV-related knowledge and behavior in which they were asked if they had heard of KS and to describe the cause of KS.

**Results**—Of 398 MSM interviewed, 73.0% (95% CI: 65.0% to 79.7%) had heard of KS. However, only 6.4% (95% CI: 4.4% to 9.2%) of participants correctly identified that KS is caused by KSHV or a virus other than HIV. Postgraduate education, urban residence, and concurrent HIV infection were all independently associated with greater awareness of the viral origin of KS.

**Conclusion**—Awareness of KSHV is very low overall among MSM and only somewhat higher, but still unacceptably low, among HIV-infected MSM. Significant efforts are needed to increase awareness of KSHV as a sexually transmitted infection in this subpopulation

#### Keywords

men who have sex with men; homosexuality; male; Kaposi sarcoma; herpesvirus 8; human; herpesvirus; Kaposi's sarcoma-associated; sampling studies

Kaposi's sarcoma (KS) was the initial malignancy described [1] and remains the most common malignancy worldwide associated with HIV disease [2,3]. Among HIV-infected individuals, the disproportionate incidence of KS among men who have sex with men (MSM) relative to other risk groups led to the hypothesis that a sexually transmitted infection, besides HIV, was causally responsible for KS [4]. This speculation was later proven when Kaposi's sarcoma-associated herpesvirus (KSHV), also known as human herpesvirus 8 (HHV-8), was discovered [5] and etiologically linked to KS [6–8]. Amidst the large and growing body of scientific knowledge about KSHV is that in the U.S. and Northern Europe KSHV is sexually transmitted and predominantly found in MSM [8,9]. Specifically, KSHV seroprevalence is 30–60% in HIV-infected MSM, 20–30% in HIV-uninfected MSM, and less than 10% in other populations [8–13].

Potential conflicts of interest: All authors: no conflicts.

Corresponding Author: Jeffrey N. Martin, M.D., M.P.H., University of California, San Francisco, 185 Berry Street, Suite 5700, San Francisco, CA 94107, Phone: 415-514-8010, Email: martin@psg.ucsf.edu.

<sup>\*</sup>Current affiliation: Oregon State University

Despite the high prevalence of KSHV infection among MSM and the threat it poses for development of KS, little is known about the awareness of KSHV in this group. Gauging the level of awareness of KSHV is important, for example, in crafting prevention messages to MSM as to how they might avoid infection with this virus. To address our limited knowledge in this area, we determined the magnitude and determinants of awareness of KSHV in a population-based sample of MSM.

Our population-based sample was drawn from the California Health Interview Survey (CHIS), a random digit-dial survey that interviews more than 50,000 adults every other year [14]. The CHIS Men who Have Sex with Men (MSM) Follow-up Study was conducted in 2002 to investigate HIV-related knowledge and behavior among MSM [15]. Of the 875 men aged 18–64 who self-identified as gay (593) or bisexual (282) in the CHIS, 741 (84.7%) agreed to be re-contacted for the Follow-up Study. No statistical differences were found between the men who agreed to participate and those who did not. Among the 741 participants who gave consent for re-contact, 193 (26.0%) could not be reached and 114 (15.4%) were excluded because they had not had sex with a man in the past 10 years. Of the remaining 434 (58.6%), 398 (91.7%) completed the follow-up interview.

All interviews were conducted in English or Spanish. Age, race/ethnicity, education, annual income, HIV infection status and area of residence were collected for all participants. In regards to this particular report, all participants were asked if they had heard of "Kaposi's sarcoma or KS." If a participant answered "yes", then he was asked to name the cause of KS. If he responded "a virus", then he was asked to identify the specific name of the virus.

In terms of statistical analysis, sampling weights designed for CHIS accounted for probability of selection, non-response, and undercoverage [16]. All analyses were conducted using the SVY procedures in Stata (College Station, TX), which adjust point estimates and standard errors to accommodate for the complex study design and sampling weights. Because of the sampling weights, final survey estimates, such as proportions, may differ from raw estimates. Correlates of the awareness of KSHV were evaluated using logistic regression. Factors associated (at p < 0.05 level) with awareness of KSHV were subsequently evaluated in a multivariable logistic regression model.

Of the 398 MSM in the survey, the majority (63.3%) were between 30 and 49 years old and were of white race/ethnicity (67.4%). Approximately half (52.7%) of the men did not have a college degree, while 31.3% reported having a college degree and 16.0% reported having a postgraduate degree. Slightly under half (44.6%) of the men had an annual income > \$60,000, while 39.5% had an income between \$20,000 and \$60,000/year, and 15.9% had an income < \$20,000/year. The majority of participants resided in an urban area (65.3%); 23.3% lived in a suburban area, and 11.3% lived in a rural area. Sixteen percent reported being HIV-infected.

We found that, of the 398 survey participants, 73.0% responded that they had heard of KS (Table 1). Regarding knowledge of the cause of KS, the most common response was that it was caused by HIV (35.2%) (Table 1). Only 17 participants (2.5%) named KSHV or human herpesvirus 8 as the cause of KS, while 3 (0.5%) stated a "herpesvirus", and 19 others (3.4%) stated a virus other than HIV but were not able to provide a name. When collectively considering mention of KSHV, HHV-8, a "herpesvirus", or any virus other than HIV, 6.4% of participants (95% CI 4.4% to 9.2%) were deemed to have provided a correct response regarding awareness of a viral etiologic agent of KS.

In evaluating the determinants of a correct response regarding awareness of KSHV (among all 398 participants), multivariable logistic regression analysis identified three independent factors: HIV infection, urban residence, and holding a postgraduate degree (Table 2). Among these, persons holding a postgraduate degree had the highest absolute level of KSHV awareness

Sex Transm Dis. Author manuscript; available in PMC 2009 December 1.

(20.8%), which translated to a 10.6-fold greater odds (95% CI 3.3 to 34.2) than those without a college degree. There was no strong evidence that age, number of sexual partners, interaction with the STD-oriented health care system, income, or race/ethnicity were independently associated with awareness of KSHV.

Following the discovery of KSHV in 1994, it has been stated that there are "few instances in cancer research where our understanding of the pathogenesis of a human cancer has progressed as rapidly as it has for Kaposi's sarcoma [17]." A portion of this progress has been in the epidemiologic characterization of KSHV infection where, in the U.S. and Northern Europe, MSM have by far the highest seroprevalence [8,9]. In view of this tremendous scientific progress, we wondered how much knowledge had been translated to MSM, the sub-population most affected by the virus. In a population-based sample of MSM living in California, we found a moderate level of awareness of KS, but a very low level of awareness of KSHV. Awareness of KSHV was higher among HIV-infected MSM, the group at most risk for development of KS, but even among this sub-group awareness is still low in absolute terms.

There are several possible reasons why awareness of KSHV is so low. First, KSHV was discovered in 1994 and causally related to KS shortly thereafter, temporally coincident with the advent of combination antiretroviral therapy for HIV. The ensuing optimism associated with antiretroviral treatment and the subsequent decline in AIDS-related opportunistic infections/malignancies, such as KS, has resulted in decreased public attention to all AIDS-related complications. (Yet, KS continues to occur in HIV-infected individuals despite seemingly effective antiretroviral therapy [18].) Second, the considerable uncertainty regarding the specific route of KSHV transmission among MSM [19] has likely weakened attempts at prevention and education about the virus. For example, guidelines state that the routes of KSHV transmission appear to be "oral, (via) semen, and through blood via needle sharing" and that patients should be counseled that "kissing and sexual intercourse with persons who have high risk for being infected with HHV-8 (e.g., persons who have KS or who are HIV-infected) might lead to acquisition of the agent that causes KS [20]." This guideline may be so broad that it discourages attempts at community education and prevention.

Our finding of low KSHV awareness among MSM is consistent with other data on how often MSM volitionally exchange saliva among each other. Saliva is the body fluid that harbors KSHV most often [21]. If MSM were highly aware of KSHV and in which body fluids it resides, we might expect to see a low prevalence of explicit and avoidable saliva-exchanging behavior. Yet, among MSM in a population-based cohort, 47% used saliva as a lubricant in the act of fingering/fisting in the prior 6 months [22]; in a different study of sexual practices among MSM, 39% reported use of saliva as a lubricant for anal sex sometime in their lifetime [23].

A potential, but still theoretical, benefit of increasing the level of awareness of KSHV among MSM is that it might reduce KSHV transmission. That KSHV is not ubiquitous among MSM suggests one of three potential explanations: a) there is some type of natural genetic-based immunity that protects about half of the population; b) KSHV infection is truly ubiquitous but serologic detection is insensitive (i.e., there is a large segment of serosilent infection); or c) there are some discrete behavioral determinants (e.g., certain saliva-exchanging sexual practices such as those which involve saliva coming in contact with the rectum) as to why some MSM are KSHV-infected and others not. Because we are not aware of any data supporting the first two explanations (or any analogy among the other human herpesviruses), we hypothesize that behavioral determinants are the main explanation. Accordingly, if certain saliva-exchanging practices are operative in spreading KSHV among MSM and if greater awareness of KSHV led to a reduction in these practices, it is conceivable that KSHV transmission could be reduced. We acknowledge, however, that even if certain saliva-exchanging acts can be

Sex Transm Dis. Author manuscript; available in PMC 2009 December 1.

identified that transmit KSHV, it may not be possible, depending upon the specific act, to reduce practice of the act in the community.

Awareness of KSHV among MSM appears to be lower than awareness of other oncogenic viruses of similar clinical importance. For example, in one report, 44.8% of MSM had heard of human papillomavirus (HPV) [24], compared to the less than 10% awareness of KSHV in our study. This finding, coupled with sub-optimal hepatitis B vaccination coverage, demonstrate need for a broader public health message educating MSM about all oncogenic viruses that affect them.

The major strength of our work is the generalizability afforded by our population-based sample with a broad representation of racial, socioeconomic, and residential status. A potential limitation of the study is that a fraction of potential participants either did not agree to be recontacted for this survey or could not be contacted despite their prior willingness. Whether these non-participators would have had more or less awareness of KSHV than the study sample is not known. That the sample was limited to Californians is also a potential limitation, but there is little reason to believe that the level of KSHV awareness we observed underestimates nationwide trends. Finally, another potential limitation is that we used self-identification as gay or bisexual as an inclusion criterion; thus, we might be missing other men who truly have sex with men but do not consider themselves to be either gay or bisexual. Again, it is not known whether those who do not self-identify as gay/bisexual would have had more or less awareness of KSHV than the study sample.

In conclusion, MSM have inadequate awareness about KSHV. MSM who are HIV-infected, more educated, and urban-dwelling are more aware, but for each of these groups awareness is still unacceptably low in absolute terms. The scientific agenda for KSHV must now expand to include determining how KSHV is spread among MSM and educating these men about KSHV, their notable risk for KSHV infection, and how -- if possible -- to avoid it.

#### Acknowledgements

Supported by the National Institutes of Health (U01 CA078124 and R01 CA119903) and the California Department of Health Services, Office of AIDS (Cooperative agreement 01-16085).

#### References

- Hymes KB, Cheung T, Greene JB, et al. Kaposi's sarcoma in homosexual men a report of eight cases. Lancet 1981;2(8247):598–600. [PubMed: 6116083]
- Mbulaiteye SM, Katabira ET, Wabinga H, et al. Spectrum of cancers among HIV-infected persons in Africa: the Uganda AIDS-Cancer Registry Match Study. International Journal of Cancer 2006;118(4): 985–990.
- International Collaboration on HIV and Cancer. Highly active antiretroviral therapy and incidence of cancer in human immunodeficiency virus-infected adults. Journal of the National Cancer Institute 2000 Nov 15;92(22):1823–1830. [PubMed: 11078759]
- Beral V, Peterman TA, Berkelman RL, et al. Kaposi's sarcoma among persons with AIDS: a sexually transmitted infection? Lancet 1990;335(8682):123–128. [PubMed: 1967430]
- Chang Y, Cesarman E, Pessin MS, et al. Identification of herpesvirus-like DNA sequences in AIDSassociated Kaposi's sarcoma. Science 1994;266(5192):1865–1869. [PubMed: 7997879]
- Whitby D, Howard MR, Tenant-Flowers M, et al. Detection of Kaposi's sarcoma-associated herpesvirus in peripheral blood HIV-infected individuals and progression to Kaposi's sarcoma. Lancet 1995;346(8978):799–802. [PubMed: 7674745]
- Gao SJ, Kingsley L, Hoover DR, et al. Seroconversion to antibodies against Kaposi's sarcomaassociated herpesvirus-related latent nuclear antigens before the development of Kaposi's sarcoma. New England Journal of Medicine 1996;335(4):233–241. [PubMed: 8657239]

Sex Transm Dis. Author manuscript; available in PMC 2009 December 1.

- Martin JN, Ganem D, Osmond DH, et al. Sexual transmission and the natural history of human herpesvirus 8 infection. New England Journal of Medicine 1998;338:948–954. [PubMed: 9521982]
- 9. Kedes DH, Opersklaski E, Busch M, et al. The seroepidemiology of human herpesvirus 8 (Kaposi's sarcoma-associated herpesvirus): distribution of infection in Kaposi's sarcoma risk groups and evidence for sexual transmission. Nature (Medicine) 1996;2(8):918–924.
- Kedes DH, Ganem D, Ameli N, et al. The prevalence of serum antibody to human herpesvirus 8 (Kaposi's sarcoma-associated herpesvirus) among HIV-seropositive and high-risk HIV-seronegative women. Journal of the American Medical Association 1997;277(6):478–481. [PubMed: 9020272]
- Dukers NH, Renwick N, Prins M, et al. Risk factors for human herpesvirus 8 seropositivity and seroconversion in a cohort of homosexual men. American Journal of Epidemiology 2000;151(3): 213–224. [PubMed: 10670545]
- Smith NA, Sabin CA, Gopal R, et al. Serologic evidence of human herpesvirus 8 transmission by homosexual but not heterosexual sex. Journal of Infectious Diseases 1999;180(3):600–606. [PubMed: 10438345]
- 13. O'Brien T, Kedes D, Ganem D, et al. Evidence for concurrent epidemics of human herpesvirus 8 and human immunodeficiency virus type 1 in US homosexual men: rates, risk factors, and relationship to Kaposi's sarcoma. Journal of Infectious Diseases 1999;180(4):1010–1017. [PubMed: 10479125]
- 14. *California Health Interview Survey, Los Angeles, CA.* 2001, Center for Health Policy Research, University of California.
- 15. Xia Q, Osmond DH, Tholandi M, et al. HIV prevalence and sexual risk behaviors among men who have sex with men. JAIDS 2006;41:238–245. [PubMed: 16394858]
- 16. *CHIS 2001 methodology series: Report 5 Weighting and variance estimation.* 2002, Center for Health Policy Research.
- 17. Beral V, Newton R, Sitas F. Human herpesvirus 8 and cancer. Journal of the National Cancer Institute 1999;91(17):1440–1441. [PubMed: 10469737]
- Maurer T, Ponte M, Leslie K. HIV-associated Kaposi's sarcoma with a high CD4 count and a low viral load. New England Journal of Medicine 2007;357(13):1352–1353. [PubMed: 17898112]
- Martin JN, Osmond DH. Invited Commentary: determining specific sexual practices associated with Human herpes virus 8 transmission. American Journal of Epidemiology 2000;151(3):225–229. [PubMed: 10670546]
- 20. Morbidity and Mortality Weekly Report. 51. 2002. Guidelines for preventing opportunistic infections among HIV-infected persons: Recommendations of the U.S. Public Health Service and the Infectious Diseases Society of America; p. 1-46.
- Pauk J, Huang ML, Brodie SJ, et al. Mucosal shedding of human herpesvirus 8 in men. New England Journal of Medicine 2000;343(19):1369–1377. [PubMed: 11070101]
- Butler, L.; Osmond, DH.; Graves Jones, A., et al. Use of saliva as a lubricant in anal sexual practices among homosexual men; Seventh International AIDS Malignancy Conference; Bethesda, Maryland. 2003.
- 23. Casper C, Carrell D, Miller KG, et al. HIV serodiscordant sex partners and the prevalence of human herpesvirus 8 infection among HIV negative men who have sex with men: baseline data from the EXPLORE study. Sexually Transmitted Infections 2006;82(3):229–235. [PubMed: 16731675]
- Pitts MK, Fox C, Willis J, et al. What do gay men know about human papillomavirus? Australian gay men's knowledge and experience of anal cancer screening and human papillomavirus. Sexually Transmitted Diseases 2007;34(3):170–173. [PubMed: 16837830]

Table 1	
Awareness of KS and KSHV	in a probability sample of MSM in California.

Question	Number of Men Responding (N = 398)	Percentage of Men Responding (95% CI)	
Awareness of KS			
Have not heard of KS	70	27.0 (20.3 to 35.0)	
Heard of KS	328	73.0 (65.0 to 79.7)	
Knowledge of the cause of KS			
HIV or HIV/AIDS	166	35.2 (29.0 to 41.9)	
Don't know	118	30.4 (23.8 to 37.8)	
Sexual contact	3	0.6 (0.2 to 2.0)	
Poppers	1	0.1 (0.02 to 1.0)	
"Hepatitis retrovirus"	1	0.3 (0.04 to 2.1)	
A virus other than HIV <sup>*</sup>	19	3.4 (2.0 to 5.7)	
KSHV or HHV-8 <sup>*</sup>	17	2.5 (1.4 to 4.4)	
"A herpes virus"*	3	0.5 (0.1 to 1.8)	

Any of these responses: 6.4% (4.4 to 9.2)

 Table 2

 Factors associated with awareness of KSHV as the causal agent of KS in a probability sample of MSM in California.

			Odds Ratio (95% CI)
Factor	Percentage of men correctly identifying viral origin of KS <sup>*</sup>	Unadjusted	$\mathbf{Adjusted}^{T}$
HIV infection status			
Uninfected/don't know	5.0	1.0	1.0
Infected	13.8	3.1 (1.3 to 7.4)	3.6 (1.3 to 9.9)
Age, in years			
< 30	2.7	1.0	1.0
30 to 49	6.7	2.6 (0.7 to 9.8)	0.8 (0.2 to 3.7)
$\geq$ 50	12.0	4.8 (1.2 to 20.3)	1.6 (0.3 to 9.9)
Place of residence			
Rural	1.7	1.0	1.0
Urban	8.4	5.5 (1.1 to 27.0)	6.4 (1.1 to 36.5)
Suburban	3.2	2.0 (0.3 to 12.4)	2.9 (0.4 to 19.2)
Education			
< College graduate	2.6	1.0	1.0
College graduate	5.6	2.2 (0.8 to 6.4)	2.3 (0.8 to 6.9)
Postgraduate degree	20.8	9.9 (3.5 to 28.2)	10.6 (3.3 to 34.2)
Interaction with STD care system <sup>#</sup>			
No	5.0	1.0	-
Yes	7.0	1.4 (0.6 to 3.4)	-
Sexual partners in past 12 months			
0	4.2	1.0	-
1 to 4	6.5	1.6 (0.3 to 7.8)	-
$\geq$ 5	7.1	1.7 (0.3 to 8.9)	-
Income, in dollars			
< 20,000	5.5	1.0	-
20,000 to 60,000	2.9	0.5 (0.1 to 2.3)	-
$\geq$ 60,000	9.7	1.9 (0.5 to 6.9)	-
Race/ethnicity		. ,	
Non-white	3.2	1.0	-
White	8.0	2.7 (0.9 to 8.4)	-

\* Correct responses include mention of KSHV, HHV-8, a herpesvirus, or a virus other than HIV.

 $^{\dagger}$ As obtained from a multivariable logistic regression model. All factors in column are included in model.

<sup>#</sup>Defined as being tested for a sexually transmitted disease (STD) in the past year