Appendix 15

Fuller details of the analyses relating lung cancer risk to the number of cigarettes smoked by the husband – adjusted for confounding and corrected for misclassification

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Random-effects Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	RR	RRL	RRU	CA1	CA0	CO1	CO0	NCigs
Akiba	Japan	1.2243	0.5745	2.4599	22.6080	16.4203	65.3375	58.1006	1.0000
Akiba	Japan	1.3197	0.6034	2.8781	17.0098	16.4203	45.6050	58.1006	2.1350
Akiba	Japan	1.6681	0.4255	6.8217	3.4794	16.4203	7.3802	58.1006	3.8390
Boffetta 2	7 countries	0.8827	0.4291	1.7249	22.2825	24.8774	49.3035	48.5890	1.8175
Boffetta	7 countries	0.9160	0.7011	1.2073	158.4235	163.6518	302.4692	286.2119	1.3380
Boffetta	7 countries	0.4811	0.2836	0.7941	21.4699	163.6518	78.0559	286.2119	2.5880
Boffetta	7 countries	1.0104	0.6138	1.6684	28.4428	163.6518	49.2304	286.2119	4.3060
Brownson 1	USA	1.5973	0.3437	7.0771	3.3190	11.8737	5.5973	31.9844	2.3157
Brownson 2	USA	0.7791	0.6054	0.9626	177.6701	216.6056	547.0460	519.6023	2.3157
Buffler	USA	0.5757	0.2193	1.5260	20.7948	7.0478	111.3676	21.7311	2.3157
Butler	USA	1.8748	0.4333	8.1692	2.6427	5.4647	11104.6809	43052.0000	2.3157
Cardenas	USA	1.0351	0.4574	2.1293	8.4607	27.8899	10856.4646	37043.2247	1.0040
Cardenas	USA	1.0460	0.5726	2.0436	14.3718	27.8899	18248.9066	37043.2247	2.2670
Cardenas	USA	1.4630	0.6822	3.1287	8.6765	27.8899	7876.8487	37043.2247	4.3140
Chan	China	0.5770	0.3028	1.0926	21.9493	42.0781	50.3072	55.6428	1.8175
Choi	Korea	1.5122	0.8433	2.6950	44.6429	25.4953	82.2954	71.0733	1.8175
Correa	USA	1.6775	0.5025	5.5573	7.4822	5.2646	35.3434	41.7168	2.3157
De Waard	Netherlands	2.3925	0.7481	7.6383	16.6844	3.7665	110.2753	59.5596	1.8175
Du	China	0.6081	0.2831	1.3060	11.9739	25.3434	60.9088	78.3930	1.0000
Du	China	1.1433	0.5688	2.2980	18.3416	25.3434	49.6254	78.3930	2.6030
Enstrom	USA	0.8498	0.5904	1.2150	91.6780	43.0465	14113.8866	5631.3381	2.3157
Fontham	USA	1.0807	0.8534	1.3683	281.0684	165.3575	525.2877	333.9623	2.3157
Gao	China	1.1514	0.7520	1.7496	146.2006	45.5462	215.1368	77.1721	1.8175
Garfinkel 1	USA	1.2132	0.8185	1.8121	38.2200	66.7380	37744.1254	79956.4373	1.0040
Garfinkel 1	USA	0.9703	0.6636	1.4168	44.4860	66.7380	54929.7729	79956.4373	2.7520
Garfinkel 2	USA	0.8004	0.3777	1.6963	10.8139	42.7707	42.4804	134.4868	1.0040
Garfinkel 2	USA	0.9650	0.5602	1.6624	28.2910	42.7707	92.1859	134.4868	2.2670
Garfinkel 2	USA	1.6032	0.8764	2.9330	23.5347	42.7707	46.1574	134.4868	4.3140
Geng	China	1.3540	0.4589	4.0153	8.0250	12.2029	14.9131	30.7056	0.4850
Geng	China	1.7864	0.5051	6.3105	5.6911	12.2029	8.0161	30.7056	1.2730
Geng	China	2.2822	0.8651	6.0417	14.5195	12.2029	16.0083	30.7056	2.6030
Gorlova	USA	1.0258	0.5446	1.9329	55.9792	27.8120	58.5452	29.8377	2.3157
Hirayama	Japan	1.2656	0.8542	1.8834	95.5493	33.0097	41848.8825	18296.9630	0.8600
Hirayama	Japan	1.3076	0.8229	2.0825	38.5747	33.0097	16351.2536	18296.9630	2.6030
Hole	Scotland	1.2784	0.1146	14.4496	2.2765	0.9197	735.0262	379.6062	0.2080
Hole	Scotland	2.2320	0.1643	30.0296	1.4615	0.9197	270.2712	379.6062	2.4090
Humble 1	USA	1.4363	0.3160	6.8716	4.3305	6.2408	6.5629	13.5844	1.5820
Humble 1	USA	0.7396	0.0898	6.7038	1.3717	6.2408	4.0371	13.5844	3.3920
IARC: Kreuzer	Germany	0.7041	0.4319	1.1658	45.6199	44.8990	122.2473	84.7110	1.8175
Inoue	Japan	2.4938	0.2949	6.5115	6.5682	3.1852	11.6304	14.0652	1.0000
Inoue	Japan	2.8284	0.7376	14.5700	8.4486	3.1852	13.1904	14.0652	2.6030
Janerich	USA	0.5698	0.3414	0.9537	54.5758	64.5956	71.1712	48.0004	2.3157
Jee	Korea	1.9047	1.0403	3.7404	52.5738	11.4150	75210.1141	31103.7062	1.0000
Jee	Korea	1.3210	0.6053	2.9599	13.0874	11.4150	26994.7380	31103.7062	2.6030
Johnson	Canada	0.9418	0.4387	2.0021	24.4405	10.4188	266.5997	107.0327	1.8175
Kabat 1	USA	0.6449	0.1891	2.1585	10.1963	10.5407	12.9606	8.6405	2.3157
Kabat 2	USA	0.6847	0.3309	1.4248	14.4550	29.5307	48.0479	67.2057	1.6540
Kabat 2	USA	0.7128	0.2900	1.7578	8.0758	29.5307	25.7822	67.2057	3.6380
Kalandidi	Greece	1.4757	0.7562	2.8800	33.0561	26.8524	37.5118	44.9678	1.3380
Kalandidi	Greece	1.6075	0.7450	3.4682	20.7493	26.8524	21.6161	44.9678	2.9020
Kalandidi	Greece	1.3185	0.4478	3.8822	7.3224	26.8524	9.3002	44.9678	5.3350
Koo	China	2.2242	0.8581	5.7838	12.3961	22.3428	11.2047	44.9192	0.7080
Koo	China	1.5495	0.6808	3.5383	16.1317	22.3428	20.9305	44.9192	1.7670
Koo	China	0.9657	0.3364	2.7289	6.7841	22.3428	14.1236	44.9192	3.1830
Lagarde	Sweden	0.9830	0.6962	1.3929	65.1667	108.1674	240.2760	392.0293	1.8175
Lam T	China	2.0811	1.0713	4.0246	21.2413	77.3621	20.9598	158.8645	0.7080
Lam T	China	1.6476	1.0190	2.6584	44.3633	77.3621	55.2935	158.8645	1.7670

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Random-effects Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	RR	RRL	RRU	CA1	CA0	CO1	CO0	NCigs
Lam T	China	1.6800	0.8091	3.5107	15.0297	77.3621	18.3709	158.8645	3.1830
Lam W	China	1.7958	0.9144	3.5394	28.7887	20.0360	52.0841	65.0950	1.8175
Layard	USA	0.5476	0.1992	1.5054	4.5765	23.0300	314.6986	867.1566	0.8190
Layard	USA	0.4629	0.1814	1.1812	5.5049	23.0300	447.7473	867.1566	2.6640
Lee	England	0.7937	0.2629	2.4021	12.8999	7.5846	28.8047	13.4416	1.8175
Lee C-H	Taiwan	1.7413	1.1980	2.5302	137.5823	59.9618	186.4875	141.5241	1.8175
Liu Q	China	0.6483	0.2027	2.1417	5.4035	12.1009	17.9456	26.0547	1.0000
Liu Q	China	2.3750	0.8631	6.8071	13.4355	12.1009	12.1801	26.0547	2.6030
Liu Z	China	0.6395	0.2344	1.7308	27.9251	6.4503	112.0352	16.5505	1.8175
Malats 7	Europe+Brazil	1.3514	0.6777	2.6839	31.9992	44.5386	22.0128	41.4064	1.8175
McGhee	China	1.1724	0.7617	1.8168	59.0307	65.4582	104.3203	135.6164	1.8175
Nishino	Japan	1.7092	0.6014	4.6209	8.1458	6.7607	18984.6435	26931.6602	1.8175
Ohno	Japan	0.8915	0.5893	1.3464	121.9982	51.5315	217.1462	81.7694	1.8175
Pershagen	Sweden	1.0657	0.6040	1.9194	30.6690	26.5218	125.0037	115.1986	1.8175
Rapiti	India	1.0356	0.3938	2.7425	9.7309	20.5828	15.5262	34.0101	1.8175
Rylander	Sweden	1.1265	0.3955	3.2152	5.2239	18.5323	30.9726	123.7757	1.8175
Schwartz	USA	0.8950	0.5683	1.4090	98.2418	63.7042	98.0407	56.8996	2.3157
Seow	Singapore	1.1332	0.7930	1.6193	74.1148	78.9212	260.8539	314.7664	1.8175
Shen	China	0.6194	0.1978	1.9956	11.1295	13.9828	12.2641	9.5445	0.4850
Shen	China	0.9253	0.2736	3.0706	11.5332	13.9828	8.5078	9.5445	1.2730
Shen	China	0.5406	0.2032	1.4463	24.0914	13.9828	30.4212	9.5445	2.6030
Shimizu	Japan	0.9488	0.5448	1.6502	42.7841	35.6817	79.3342	62.7763	1.8175
Sobue	Japan	0.9939	0.6679	1.4725	63.8258	54.6271	324.9230	276.3902	1.8175
Speize	USA	1.2129	0.1982	6.2355	17.7778	1.3930	189387.4950	18000.0001	2.3157
Stockwell	USA	1.3666	0.6460	2.7100	37.4784	17.9515	49.1731	32.1867	2.3157
Sun	China	0.9682	0.6436	1.4633	109.6797	78.2982	111.1534	76.8271	1.8175
Svensson	Sweden	1.1740	0.4252	3.2419	14.2516	6.3903	69.2020	36.4291	1.8175
Trichopoulos	Greece	0.5557	0.1196	2.5822	1.9964	24.8229	15.6397	108.0606	0.7080
Trichopoulos	Greece	2.3682	1.1930	4.7009	21.1990	24.8229	38.9686	108.0606	1.7670
Trichopoulos	Greece	3.6754	1.2102	11.1626	6.7874	24.8229	8.0392	108.0606	2.5880
Trichopoulos	Greece	1.6422	0.6088	4.4297	6.6599	24.8229	17.6544	108.0606	4.3060
Vineis	Western Europe	0.9687	0.4923	1.9209	19.4033	14.4566	26955.3348	19455.3656	1.8175
Wang L	China	0.9999	0.5801	1.6571	120.9369	25.1228	246.1107	51.1209	1.8175
Wang S	China	2.3678	1.1689	4.8149	63.0823	15.0977	57.2066	32.4182	1.8175
Wang T	China	0.3336	0.1035	1.1197	4.1573	41.3005	12.4810	41.3572	0.4850
Wang T	China	1.1898	0.6306	2.2331	39.3062	41.3005	33.0823	41.3572	1.2730
Wang T	China	1.0812	0.5479	2.1263	29.1814	41.3005	27.0257	41.3572	2.6030
Wu	USA	1.0462	0.4060	3.0890	14.4212	9.1903	27.8279	18.5531	2.3157
Wu-Williams	China	0.5572	0.4637	0.7378	200.4976	294.5987	392.9901	321.7532	1.8175
Zaridze	Russia	1.5965	1.0471	2.4264	60.4727	81.5419	85.3327	183.6968	0.7080
Zaridze	Russia	1.1898	0.7375	1.9284	36.0290	81.5419	68.2201	183.6968	2.2930
Zatloukal	Czech Republic	0.2425	0.0558	1.0467	1.9370	37.8270	73.3741	347.4617	1.8175
Zheng	China	2.2987	0.9064	5.8462	46.9414	5.8178	136.8707	38.9939	1.8175
Zhong	China	1.3279	0.8422	2.1153	78.7277	60.4097	76.3714	77.8183	0.7080
Zhong	China	0.7888	0.5083	1.2693	64.2590	60.4097	104.9445	77.8183	1.7670
Zhong	China	1.1039	0.5193	2.1791	17.7742	60.4097	20.7416	77.8183	3.1830

NOTES: RR,RL,RU RELATIVE RISKS AND LOWER AND UPPER CONFIDENCE INTERVALS

CA1,CA0 NUMBERS (OR PSEUDO-NUMBERS) OF EXPOSED AND UNEXPOSED CASES

CO1,CO0 NUMBERS (OR PSEUDO-NUMBERS) OF EXPOSED AND UNEXPOSED CONTROLS

NCIGS MIDPOINT OF NUMBER OF CIGARETTES SMOKED BY HUSBAND (IN UNITS OF 10)

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Random-effects Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	Beta	St.Err. Beta	Z	Weight	ETSse	FRU	VEG	FAT	EDU	TEA
Akiba	Japan	0.1270	0.1468	0.8651	46.4032	0.9896	0	0	0	0	0
Boffetta 2	7 countries	-0.0686	0.1953	-0.3513	26.2178	0.9130	0	0	0	0	0
Boffetta	7 countries	-0.0579	0.0531	-1.0904	354.6590	1.1768	0	0	0	0	0
Brownson 1	USA	0.2022	0.3332	0.6068	9.0072	0.8345	0	0	0	1	0
Brownson 2	USA	-0.1078	0.0511	-2.1096	382.9642	1.1579	0	0	0	0	0
Buffler	USA	-0.2384	0.2137	-1.1156	21.8973	0.8581	0	0	0	0	0
Butler	USA	0.2714	0.3235	0.8389	9.5555	0.9215	0	0	0	0	0
Cardenas	USA	0.0742	0.0845	0.8781	140.0511	1.4556	0	1	1	2	0
Chan	China	-0.3026	0.1801	-1.6802	30.8299	0.9109	0	0	0	0	0
Choi	Korea	0.2276	0.1631	1.3955	37.5917	0.9091	0	0	0	0	0
Correa	USA	0.2234	0.2647	0.8440	14.2722	1.1582	0	0	0	0	0
De Waard	Netherlands	0.4800	0.3261	1.4719	9.4037	0.8696	0	0	0	0	0
Du	China	0.0570	0.1368	0.4167	53.4352	1.0879	0	0	0	0	0
Enstrom	USA	-0.0703	0.0795	-0.8843	158.2216	1.0456	1	0	0	2	0
Fontham	USA	0.0335	0.0520	0.6442	369.8225	1.1293	1	1	0	2	0
Gao	China	0.0776	0.1185	0.6549	71.2137	0.8022	0	0	0	2	0
Garfinkel 1	USA	-0.0138	0.0703	-0.1963	202.3435	1.1963	0	0	0	0	0
Garfinkel 2	USA	0.0979	0.0689	1.4209	210.6500	1.5345	0	0	0	0	0
Geng	China	0.3058	0.1831	1.6701	29.8279	1.0478	0	0	0	0	0
Gorlova	USA	0.0110	0.1395	0.0789	51.3868	1.1005	0	0	0	1	0
Hirayama	Japan	0.0781	0.0867	0.9008	133.0337	0.8998	0	0	0	0	0
Hole	Scotland	0.2824	0.4379	0.6449	5.2149	1.0530	0	0	0	0	0
Humble 1	USA	-0.0149	0.3014	-0.0494	11.0081	1.2885	0	0	0	0	0
IARC: Kreuzer	Germany	-0.1931	0.1394	-1.3852	51.4606	0.8957	0	0	0	0	0
Inoue	Japan	0.3216	0.2758	1.1661	13.1465	1.1196	0	0	0	0	0
Janerich	USA	-0.2429	0.1132	-2.1458	78.0382	1.1397	0	0	0	0	0
Jee	Korea	0.0267	0.1509	0.1769	43.9159	0.9132	0	1	0	3	0
Johnson	Canada	-0.0330	0.2131	-0.1549	22.0208	0.8225	1	1	0	2	0
Kabat 1	USA	-0.1894	0.2682	-0.7062	13.9022	1.1579	0	0	0	0	0
Kabat 2	USA	-0.1142	0.1214	-0.9407	67.8520	1.3646	0	0	0	2	0
Kalandidi	Greece	0.0807	0.0916	0.8810	119.1816	1.5584	0	0	0	0	0
Koo	China	0.0293	0.1532	0.1913	42.6071	1.1884	0	0	0	2	0
Lagarde	Sweden	-0.0095	0.0973	-0.0976	105.6268	0.8828	0	0	0	3	0
Lam T	China	0.2048	0.0984	2.0813	103.2785	1.0055	0	0	0	0	0
Lam W	China	0.3221	0.1900	1.6953	27.7008	0.9063	0	0	0	0	0
Layard	USA	-0.3026	0.1784	-1.6962	31.4203	1.1466	0	0	0	0	0
Lee	England	-0.1271	0.3105	-0.4093	10.3723	0.8544	0	0	0	0	0
Lee C-H	Taiwan	0.3052	0.1049	2.9094	90.8760	0.9015	0	0	0	2	0
Liu Q	China	0.3460	0.2022	1.7112	24.4589	1.0561	0	0	0	2	0
Liu Z	China	-0.2459	0.2807	-0.8760	12.6916	0.6106	0	0	0	0	0
Malats	7 Europe+Brazil	0.1657	0.1932	0.8577	26.7908	0.8717	0	0	0	0	0
McGhee	China	0.0875	0.1220	0.7172	67.1862	0.9025	0	0	0	2	0
Nishino	Japan	0.2949	0.2862	1.0304	12.2085	0.8950	1	1	1	0	0
Ohno	Japan	-0.0632	0.1160	-0.5448	74.3163	0.8114	0	0	0	0	0
Pershagen	Sweden	0.0350	0.1623	0.2157	37.9632	0.9097	0	0	0	0	0
Rapiti	India	0.0192	0.2724	0.0705	13.4768	0.8504	0	0	0	0	0
Rylander	Sweden	0.0655	0.2941	0.2227	11.5614	0.7290	0	0	0	0	0
Schwartz	USA	-0.0479	0.1000	-0.4790	100.0000	1.1194	0	0	0	0	0
Seow	Singapore	0.0688	0.1002	0.6866	99.6012	0.9054	0	0	0	0	0
Shen	China	-0.1830	0.1682	-1.0880	35.3466	1.0903	0	0	0	0	0
Shimizu	Japan	-0.0289	0.1556	-0.1857	41.3029	0.9053	0	0	0	0	0
Sobue	Japan	-0.0034	0.1110	-0.0306	81.1622	0.9064	0	0	0	2	0
Speize	USA	0.0834	0.3799	0.2195	6.9289	0.6375	0	0	0	0	0
Stockwell	USA	0.1349	0.1580	0.8538	40.0577	1.1381	0	0	0	2	0
Sun	China	-0.0178	0.1153	-0.1544	75.2214	0.8954	0	0	0	2	0
Svensson	Sweden	0.0883	0.2851	0.3097	12.3028	0.8673	0	0	0	0	0
Trichopoulos	Greece	0.2307	0.1069	2.1581	87.5074	1.3389	0	0	0	0	0

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Random-effects Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	Beta	St.Err. Beta	Z	Weight	ETSse	FRU	VEG	FAT	EDU	TEA
Vineis	Western Europe	-0.0175	0.1911	-0.0916	27.3828	0.8968	1	1	1	2	0
Wang L	China	-0.0001	0.1473	-0.0007	46.0887	0.6870	0	0	0	1	0
Wang S	China	0.4742	0.1987	2.3865	25.3282	0.8780	0	0	0	0	0
Wang T	China	0.0736	0.1308	0.5627	58.4500	1.0379	0	0	0	0	0
Wu	USA	0.0195	0.2236	0.0872	20.0012	1.1449	0	0	0	0	0
Wu-Williams	China	-0.3218	0.0652	-4.9356	235.2366	0.9047	0	0	0	2	0
Zaridze	Russia	0.0796	0.1069	0.7446	87.5074	0.8744	0	0	0	2	0
Zatloukal	Czech Republic	-0.7795	0.4114	-1.8947	5.9084	0.6900	0	0	0	2	0
Zheng	China	0.4580	0.2616	1.7508	14.6125	0.7567	0	0	0	0	0
Zhong	China	-0.0804	0.0971	-0.8280	106.0624	0.9343	1	0	0	1	0

NOTES: BETA SLOPE OF LOG RR ON NCIGS

SE BETA STANDARD ERROR OF BETA

Z RATIO OF BETA TO ITS STANDARD ERROR (APPROXIMATE NORMAL STATISTIC)

WEIGHT INVERSE OF THE VARIANCE OF BETA

ETSse STANDARD ERROR OF ETS EXPOSURE OVER THE CONTROL GROUPS

FRU = 1 DOSE-RESPONSE ANALYSIS ADJUSTED FOR FRUIT

VEG = 1 " " " " VEGETABLES

FAT = 1 " " " " FRUIT

EDU = 1 " " " " INCOME

= 2 " " " " EDUCATION

= 3 " " " " SOCIO-ECONOMIC STATUS

TEA = 0 NO ADJUSTMENT (OR MATCHING) FOR TEA

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
RANDOM-EFFECTS META-ANALYSES
Confounder Differences based on Random-effects Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	61.9625	(66)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0197	0.0209	N.S.	1.0199	0.9790	1.0625

SEPARATE RANDOM-EFFECTS META-ANALYSES BY REGION

NORTH AMERICA AND EUROPE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	35.5427	(36)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0113	0.0194	N.S.	0.9888	0.9519	1.0271

ASIA
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	23.6933	(29)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0646	0.0389	N.S.	1.0667	0.9883	1.1513

USA AND CANADA
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	19.8263	(20)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0240	0.0230	N.S.	0.9763	0.9333	1.0213

EUROPE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	14.7000	(15)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0181	0.0360	N.S.	1.0183	0.9490	1.0927

CHINA AND HONG KONG
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	14.7364	(17)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0506	0.0581	N.S.	1.0519	0.9386	1.1788

REST OF ASIA

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY REGION
Confounder Differences based on Random-effects Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	9.5442	(11)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0910	0.0382	+	1.0953	1.0164	1.1803

SEPARATE RANDOM-EFFECTS META-ANALYSES BY YEAR OF PUBLICATION

PUBLISHED IN 1980S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	20.2801	(24)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0845	0.0284	++	1.0882	1.0294	1.1504

PUBLISHED IN 1990S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	27.1510	(26)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0203	0.0325	N.S.	0.9799	0.9194	1.0443

PUBLISHED IN 2000S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	13.8250	(14)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0211	0.0403	N.S.	1.0213	0.9438	1.1052

SEPARATE RANDOM-EFFECTS META-ANALYSES BY NUMBER OF LUNG CANCER CASES

<100 CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	38.0266	(38)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0602	0.0333	(+)	1.0620	0.9949	1.1336

100-199 CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	12.4821	(13)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0022	0.0272	N.S.	0.9978	0.9459	1.0525

200-399 CASES

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY NUMBER OF LUNG CANCER CASES
Confounder Differences based on Random-effects Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	7.9680	(8)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0972	0.0352	+	1.1020	1.0285	1.1808

400+ CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	4.0692	(4)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.1045	0.0587	N.S.	0.9007	0.8029	1.0106

SEPARATE RANDOM-EFFECTS META-ANALYSES BY DOSE-RESPONSE OR NOT

RESULTS FOR DOSE-RESPONSE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	21.7577	(22)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0471	0.0257	(+)	1.0483	0.9968	1.1024

NO RESULTS FOR DOSE-RESPONSE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	39.4629	(43)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0004	0.0294	N.S.	1.0004	0.9444	1.0598

SEPARATE RANDOM-EFFECTS META-ANALYSES BY AGE ADJUSTMENT OR NOT

ADJUSTED (OR MATCHED) FOR AGE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	48.2125	(54)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0103	0.0198	N.S.	0.9897	0.9521	1.0289

NOT ADJUSTED (OR MATCHED) FOR AGE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	12.1959	(11)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.1773	0.0596	+	1.1939	1.0623	1.3418

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY STUDY TYPE
Confounder Differences based on Random-effects Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

CASE-CONTROL STUDIES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	53.1306	(56)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0193	0.0242	N.S.	1.0195	0.9722	1.0691

PROSPECTIVE STUDIES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	4.3343	(9)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0214	0.0368	N.S.	1.0216	0.9505	1.0980

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Unweighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	RR	RRL	RRU	CA1	CA0	CO1	CO0	NCigs
Akiba	Japan	1.2113	0.5684	2.4340	22.4342	16.5648	64.7174	57.8845	1.0000
Akiba	Japan	1.2900	0.5898	2.8136	16.8543	16.5648	45.6555	57.8845	2.1350
Akiba	Japan	1.6012	0.4083	6.5494	3.4429	16.5648	7.5138	57.8845	3.8390
Boffetta 2	7 countries	0.8649	0.4204	1.6905	22.0426	25.1170	49.3032	48.5887	1.8175
Boffetta	7 countries	0.9025	0.6907	1.1895	156.9670	165.6416	300.5283	286.2108	1.3380
Boffetta	7 countries	0.4674	0.2755	0.7716	21.3581	165.6416	78.9578	286.2108	2.5880
Boffetta	7 countries	0.9632	0.5850	1.5906	28.0197	165.6416	50.2668	286.2108	4.3060
Brownson 1	USA	1.5674	0.3371	6.9480	3.2908	11.9976	5.6326	32.1860	2.3157
Brownson 2	USA	0.7614	0.5914	0.9411	175.1836	218.5257	546.2603	518.8560	2.3157
Buffler	USA	0.5624	0.2141	1.4914	20.4676	7.1009	110.2736	21.5169	2.3157
Butler	USA	1.8450	0.4263	8.0402	2.5884	5.7093	10972.5856	44653.7238	2.3157
Cardenas	USA	1.0340	0.4570	2.1272	8.4562	27.9356	10848.2784	37057.3992	1.0040
Cardenas	USA	1.0436	0.5712	2.0389	14.3586	27.9356	18251.3261	37057.3992	2.2670
Cardenas	USA	1.4566	0.6792	3.1151	8.6712	27.9356	7896.7666	37057.3992	4.3140
Chan	China	0.5627	0.2949	1.0669	21.6166	42.4914	50.3701	55.7137	1.8175
Choi	Korea	1.4883	0.8299	2.6525	44.2379	25.6706	82.0259	70.8406	1.8175
Correa	USA	1.6473	0.4935	5.4565	7.4079	5.3080	35.2577	41.6159	2.3157
De Waard	Netherlands	2.3607	0.7381	7.5376	16.5055	3.7779	109.3531	59.0868	1.8175
Du	China	0.6018	0.2801	1.2930	11.9174	25.5821	60.6210	78.3176	1.0000
Du	China	1.1129	0.5532	2.2387	18.1061	25.5821	49.8057	78.3176	2.6030
Enstrom	USA	0.8411	0.5844	1.2027	91.0076	43.1701	14056.5975	5608.4803	2.3157
Fontham	USA	1.0696	0.8447	1.3544	279.4347	166.0954	524.2335	333.2921	2.3157
Gao	China	1.1302	0.7381	1.7173	144.6776	45.9193	213.8451	76.7071	1.8175
Garfinkel 1	USA	1.2016	0.8106	1.7948	37.9578	67.5329	37443.1738	80044.8591	1.0040
Garfinkel 1	USA	0.9450	0.6463	1.3800	44.1186	67.5329	55333.7385	80044.8591	2.7520
Garfinkel 2	USA	0.7951	0.3752	1.6852	10.7988	43.2029	42.2706	134.4699	1.0040
Garfinkel 2	USA	0.9506	0.5518	1.6379	28.1043	43.2029	92.0171	134.4699	2.2670
Garfinkel 2	USA	1.5582	0.8516	2.8511	23.2893	43.2029	46.5199	134.4699	4.3140
Geng	China	1.3488	0.4572	3.9984	8.0139	12.3160	14.8197	30.7188	0.4850
Geng	China	1.7682	0.5004	6.2412	5.6831	12.3160	8.0164	30.7188	1.2730
Geng	China	2.2350	0.8485	5.9069	14.4427	12.3160	16.1180	30.7188	2.6030
Gorlova	USA	1.0043	0.5332	1.8924	55.3846	28.1050	58.3345	29.7304	2.3157
Hirayama	Japan	1.2512	0.8446	1.8620	94.1037	33.1903	41093.8583	18134.7303	0.8600
Hirayama	Japan	1.2633	0.7951	2.0117	38.3578	33.1903	16590.2378	18134.7303	2.6030
Hole	Scotland	1.2758	0.1144	14.4208	2.2631	0.9218	727.5610	378.0527	0.2080
Hole	Scotland	2.1790	0.1603	29.3427	1.4538	0.9218	273.6225	378.0527	2.4090
Humble 1	USA	1.4130	0.3105	6.7688	4.2858	6.3210	6.5295	13.6072	1.5820
Humble 1	USA	0.7141	0.0865	6.4906	1.3562	6.3210	4.0883	13.6072	3.3920
IARC: Kreuzer	Germany	0.6887	0.4224	1.1407	45.0825	45.3536	122.1354	84.6262	1.8175
Inoue	Japan	2.4768	0.2929	6.4674	6.5403	3.2023	11.5380	13.9923	1.0000
Inoue	Japan	2.7784	0.7245	14.3141	8.3653	3.2023	13.1554	13.9923	2.6030
Janerich	USA	0.5556	0.3326	0.9305	53.7473	65.2388	71.0621	47.9240	2.3157
Jee	Korea	1.8842	1.0294	3.6988	49.4616	11.5893	86572.5057	38219.7797	1.0000
Jee	Korea	1.2842	0.5890	2.8749	12.9284	11.5893	33199.4895	38219.7797	2.6030
Johnson	Canada	0.9317	0.4340	1.9806	24.2638	10.4559	265.5318	106.6039	1.8175
Kabat 1	USA	0.6298	0.1845	2.1101	10.0604	10.6489	12.9431	8.6289	2.3157
Kabat 2	USA	0.6735	0.3252	1.4026	14.3155	29.8941	47.8899	67.3514	1.6540
Kabat 2	USA	0.6875	0.2792	1.6982	7.9645	29.8941	26.1009	67.3514	3.6380
Kalandidi	Greece	1.4616	0.7489	2.8525	32.8938	27.1264	37.2458	44.8938	1.3380
Kalandidi	Greece	1.5743	0.7297	3.3967	20.5838	27.1264	21.6382	44.8938	2.9020
Kalandidi	Greece	1.2690	0.4310	3.7364	7.2280	27.1264	9.4267	44.8938	5.3350
Koo	China	2.2087	0.8521	5.7435	12.3810	22.6214	11.1439	44.9714	0.7080
Koo	China	1.5226	0.6690	3.4771	16.0012	22.6214	20.8914	44.9714	1.7670
Koo	China	0.9357	0.3259	2.6445	6.7209	22.6214	14.2786	44.9714	3.1830
Lagarde	Sweden	0.9632	0.6819	1.3654	64.4621	109.1902	240.7169	392.7489	1.8175
Lam T	China	2.0676	1.0643	3.9987	21.2268	78.1378	20.9084	159.1371	0.7080
Lam T	China	1.6211	1.0025	2.6159	44.0364	78.1378	55.3231	159.1371	1.7670

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Unweighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	RR	RRL	RRU	CA1	CA0	CO1	CO0	NCigs
Lam T	China	1.6317	0.7857	3.4103	14.8664	78.1378	18.5551	159.1371	3.1830
Lam W	China	1.7696	0.9017	3.4853	28.6153	20.2162	52.0853	65.1147	1.8175
Layard	USA	0.5411	0.1966	1.4893	4.5493	23.4250	313.9656	874.8296	0.8190
Layard	USA	0.4455	0.1739	1.1407	5.4300	23.4250	455.2318	874.8296	2.6640
Lee	England	0.7767	0.2572	2.3518	12.7379	7.6543	28.6767	13.3837	1.8175
Lee C-H	Taiwan	1.7155	1.1803	2.4928	136.4321	60.3527	185.7708	140.9802	1.8175
Liu Q	China	0.6434	0.2011	2.1259	5.3919	12.1868	17.8806	26.0035	1.0000
Liu Q	China	2.3289	0.8460	6.6775	13.3004	12.1868	12.1859	26.0035	2.6030
Liu Z	China	0.6271	0.2299	1.6965	27.5994	6.5012	111.1445	16.4192	1.8175
Malats 7	Europe+Brazil	1.3282	0.6659	2.6382	31.7026	44.8978	22.0308	41.4403	1.8175
McGhee	China	1.1501	0.7470	1.7830	58.3787	65.9894	104.2191	135.4848	1.8175
Nishino	Japan	1.7132	0.6028	4.6314	8.1571	6.7543	18990.9635	26940.6258	1.8175
Ohno	Japan	0.8737	0.5774	1.3197	120.5968	51.9795	215.9506	81.3216	1.8175
Pershagen	Sweden	1.0455	0.5925	1.8836	30.3392	26.7416	124.7625	114.9769	1.8175
Rapiti	India	1.0155	0.3858	2.6914	9.6293	20.7705	15.5706	34.1069	1.8175
Rylander	Sweden	1.1032	0.3864	3.1564	5.1675	18.7193	31.1430	124.4566	1.8175
Schwartz	USA	0.8765	0.5565	1.3800	97.1747	64.3439	97.7832	56.7486	2.3157
Seow	Singapore	1.1121	0.7781	1.5895	73.3544	79.5928	260.7041	314.5842	1.8175
Shen	China	0.6159	0.1967	1.9842	11.1463	14.1673	12.1533	9.5135	0.4850
Shen	China	0.9114	0.2694	3.0246	11.5051	14.1673	8.4770	9.5135	1.2730
Shen	China	0.5241	0.1970	1.4023	23.7223	14.1673	30.3972	9.5135	2.6030
Shimizu	Japan	0.9299	0.5338	1.6178	42.3063	35.9960	79.1723	62.6418	1.8175
Sobue	Japan	0.9744	0.6547	1.4440	63.0830	55.0707	324.1021	275.6919	1.8175
Speize	USA	1.1886	0.1946	6.0981	17.3947	1.3990	188299.7800	17999.9985	2.3157
Stockwell	USA	1.3422	0.6345	2.6617	37.1153	18.0999	48.9826	32.0617	2.3157
Sun	China	0.9483	0.6302	1.4336	108.4092	79.0189	110.8261	76.6006	1.8175
Svensson	Sweden	1.1527	0.4175	3.1826	14.0952	6.4353	68.8350	36.2273	1.8175
Trichopoulos	Greece	0.5533	0.1190	2.5709	1.9961	25.0440	15.5727	108.0990	0.7080
Trichopoulos	Greece	2.3424	1.1800	4.6498	21.0768	25.0440	38.8383	108.0990	1.7670
Trichopoulos	Greece	3.6170	1.1909	10.9851	6.7552	25.0440	8.0614	108.0990	2.5880
Trichopoulos	Greece	1.5990	0.5927	4.3131	6.6155	25.0440	17.8581	108.0990	4.3060
Vineis	Western Europe	0.9719	0.4940	1.9271	19.4432	14.4386	26972.8589	19468.0138	1.8175
Wang L	China	0.9796	0.5683	1.6235	119.4177	25.3208	243.8844	50.6584	1.8175
Wang S	China	2.3372	1.1538	4.7526	62.6124	15.1810	56.9220	32.2558	1.8175
Wang T	China	0.3318	0.1030	1.1140	4.1609	41.7528	12.4121	41.3315	0.4850
Wang T	China	1.1739	0.6222	2.2032	39.1022	41.7528	32.9737	41.3315	1.2730
Wang T	China	1.0520	0.5331	2.0685	28.8549	41.7528	27.1528	41.3315	2.6030
Wu	USA	1.0268	0.3985	3.0316	14.2785	9.2708	27.7544	18.5028	2.3157
Wu-Williams	China	0.5432	0.4519	0.7196	197.3344	297.4432	392.7372	321.5462	1.8175
Zaridze	Russia	1.5820	1.0376	2.4044	60.1828	82.4425	84.8801	183.9484	0.7080
Zaridze	Russia	1.1552	0.7161	1.8724	35.6629	82.4425	68.8820	183.9484	2.2930
Zatloukal	Czech Republic	0.2327	0.0524	1.0262	1.8754	38.1680	73.8896	349.9032	1.8175
Zheng	China	2.2630	0.8931	5.7501	46.4656	5.8501	135.7173	38.6684	1.8175
Zhong	China	1.3193	0.8367	2.1016	78.5109	60.9146	75.9120	77.7066	0.7080
Zhong	China	0.7761	0.5002	1.2488	63.8287	60.9146	104.9132	77.7066	1.7670
Zhong	China	1.0722	0.5044	2.1163	17.6019	60.9146	20.9426	77.7066	3.1830

NOTES: RR,RL,RU RELATIVE RISKS AND LOWER AND UPPER CONFIDENCE INTERVALS

CA1,CA0 NUMBERS (OR PSEUDO-NUMBERS) OF EXPOSED AND UNEXPOSED CASES

CO1,CO0 NUMBERS (OR PSEUDO-NUMBERS) OF EXPOSED AND UNEXPOSED CONTROLS

NCIGS MIDPOINT OF NUMBER OF CIGARETTES SMOKED BY HUSBAND (IN UNITS OF 10)

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Unweighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	Beta	St.Err. Beta	Z	Weight	ETSse	FRU	VEG	FAT	EDU	TEA
Akiba	Japan	0.1164	0.1468	0.7929	46.4032	0.9896	0	0	0	0	0
Boffetta 2	7 countries	-0.0799	0.1953	-0.4091	26.2178	0.9130	0	0	0	0	0
Boffetta	7 countries	-0.0691	0.0531	-1.3013	354.6590	1.1768	0	0	0	0	0
Brownson 1	USA	0.1941	0.3333	0.5824	9.0018	0.8345	0	0	0	1	0
Brownson 2	USA	-0.1177	0.0512	-2.2988	381.4697	1.1579	0	0	0	0	0
Buffler	USA	-0.2485	0.2138	-1.1623	21.8768	0.8581	0	0	0	0	0
Butler	USA	0.2645	0.3236	0.8174	9.5496	0.9215	0	0	0	0	0
Cardenas	USA	0.0732	0.0844	0.8673	140.3832	1.4556	0	1	1	2	0
Chan	China	-0.3164	0.1805	-1.7529	30.6934	0.9109	0	0	0	0	0
Choi	Korea	0.2188	0.1631	1.3415	37.5917	0.9091	0	0	0	0	0
Correa	USA	0.2155	0.2647	0.8141	14.2722	1.1582	0	0	0	0	0
De Waard	Netherlands	0.4726	0.3261	1.4492	9.4037	0.8696	0	0	0	0	0
Du	China	0.0459	0.1370	0.3350	53.2793	1.0879	0	0	0	0	0
Enstrom	USA	-0.0747	0.0795	-0.9396	158.2216	1.0456	1	0	0	2	0
Fontham	USA	0.0291	0.0520	0.5596	369.8225	1.1293	1	1	0	2	0
Gao	China	0.0673	0.1185	0.5679	71.2137	0.8022	0	0	0	2	0
Garfinkel 1	USA	-0.0230	0.0703	-0.3272	202.3435	1.1963	0	0	0	0	0
Garfinkel 2	USA	0.0910	0.0689	1.3208	210.6500	1.5345	0	0	0	0	0
Geng	China	0.2979	0.1829	1.6288	29.8932	1.0478	0	0	0	0	0
Gorlova	USA	0.0019	0.1395	0.0136	51.3868	1.1005	0	0	0	1	0
Hirayama	Japan	0.0653	0.0868	0.7523	132.7274	0.8998	0	0	0	0	0
Hole	Scotland	0.2726	0.4389	0.6211	5.1912	1.0530	0	0	0	0	0
Humble 1	USA	-0.0249	0.3015	-0.0826	11.0008	1.2885	0	0	0	0	0
IARC: Kreuzer	Germany	-0.2052	0.1394	-1.4720	51.4606	0.8957	0	0	0	0	0
Inoue	Japan	0.3156	0.2761	1.1431	13.1180	1.1196	0	0	0	0	0
Janerich	USA	-0.2538	0.1133	-2.2401	77.9005	1.1397	0	0	0	0	0
Jee	Korea	0.0207	0.1513	0.1368	43.6840	0.9132	0	1	0	3	0
Johnson	Canada	-0.0390	0.2131	-0.1830	22.0208	0.8225	1	1	0	2	0
Kabat 1	USA	-0.1996	0.2685	-0.7434	13.8711	1.1579	0	0	0	0	0
Kabat 2	USA	-0.1244	0.1215	-1.0239	67.7404	1.3646	0	0	0	2	0
Kalandidi	Greece	0.0737	0.0916	0.8046	119.1816	1.5584	0	0	0	0	0
Koo	China	0.0203	0.1531	0.1326	42.6628	1.1884	0	0	0	2	0
Lagarde	Sweden	-0.0206	0.0975	-0.2113	105.1940	0.8828	0	0	0	3	0
Lam T	China	0.1959	0.0983	1.9929	103.4887	1.0055	0	0	0	0	0
Lam W	China	0.3140	0.1898	1.6544	27.7592	0.9063	0	0	0	0	0
Layard	USA	-0.3177	0.1790	-1.7749	31.2100	1.1466	0	0	0	0	0
Lee	England	-0.1390	0.3106	-0.4475	10.3657	0.8544	0	0	0	0	0
Lee C-H	Taiwan	0.2970	0.1049	2.8313	90.8760	0.9015	0	0	0	2	0
Liu Q	China	0.3379	0.2022	1.6711	24.4589	1.0561	0	0	0	2	0
Liu Z	China	-0.2567	0.2805	-0.9152	12.7097	0.6106	0	0	0	0	0
Malats	7 Europe+Brazil	0.1562	0.1932	0.8085	26.7908	0.8717	0	0	0	0	0
McGhee	China	0.0769	0.1221	0.6298	67.0762	0.9025	0	0	0	2	0
Nishino	Japan	0.2962	0.2862	1.0349	12.2085	0.8950	1	1	1	0	0
Ohno	Japan	-0.0743	0.1160	-0.6405	74.3163	0.8114	0	0	0	0	0
Pershagen	Sweden	0.0245	0.1624	0.1509	37.9165	0.9097	0	0	0	0	0
Rapiti	India	0.0085	0.2726	0.0312	13.4570	0.8504	0	0	0	0	0
Rylander	Sweden	0.0540	0.2948	0.1832	11.5065	0.7290	0	0	0	0	0
Schwartz	USA	-0.0569	0.1001	-0.5684	99.8003	1.1194	0	0	0	0	0
Seow	Singapore	0.0585	0.1003	0.5833	99.4027	0.9054	0	0	0	0	0
Shen	China	-0.1951	0.1685	-1.1579	35.2209	1.0903	0	0	0	0	0
Shimizu	Japan	-0.0400	0.1556	-0.2571	41.3029	0.9053	0	0	0	0	0
Sobue	Japan	-0.0143	0.1110	-0.1288	81.1622	0.9064	0	0	0	2	0
Speize	USA	0.0746	0.3795	0.1966	6.9435	0.6375	0	0	0	0	0
Stockwell	USA	0.1271	0.1580	0.8044	40.0577	1.1381	0	0	0	2	0
Sun	China	-0.0292	0.1154	-0.2530	75.0911	0.8954	0	0	0	2	0
Svensson	Sweden	0.0782	0.2851	0.2743	12.3028	0.8673	0	0	0	0	0
Trichopoulos	Greece	0.2247	0.1068	2.1039	87.6713	1.3389	0	0	0	0	0

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Unweighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	Beta	St.Err. Beta	Z	Weight	ETSse	FRU	VEG	FAT	EDU	TEA
Vineis	Western Europe	-0.0157	0.1911	-0.0822	27.3828	0.8968	1	1	1	2	0
Wang L	China	-0.0113	0.1473	-0.0767	46.0887	0.6870	0	0	0	1	0
Wang S	China	0.4671	0.1987	2.3508	25.3282	0.8780	0	0	0	0	0
Wang T	China	0.0629	0.1308	0.4809	58.4500	1.0379	0	0	0	0	0
Wu	USA	0.0114	0.2235	0.0510	20.0191	1.1449	0	0	0	0	0
Wu-Williams	China	-0.3358	0.0653	-5.1424	234.5166	0.9047	0	0	0	2	0
Zaridze	Russia	0.0673	0.1069	0.6296	87.5074	0.8744	0	0	0	2	0
Zatloukal	Czech Republic	-0.8023	0.4175	-1.9217	5.7370	0.6900	0	0	0	2	0
Zheng	China	0.4494	0.2614	1.7192	14.6349	0.7567	0	0	0	0	0
Zhong	China	-0.0891	0.0971	-0.9176	106.0624	0.9343	1	0	0	1	0

NOTES: BETA SLOPE OF LOG RR ON NCIGS

SE BETA STANDARD ERROR OF BETA

Z RATIO OF BETA TO ITS STANDARD ERROR (APPROXIMATE NORMAL STATISTIC)

WEIGHT INVERSE OF THE VARIANCE OF BETA

ETSse STANDARD ERROR OF ETS EXPOSURE OVER THE CONTROL GROUPS

FRU = 1 DOSE-RESPONSE ANALYSIS ADJUSTED FOR FRUIT

VEG = 1 " " " " VEGETABLES

FAT = 1 " " " " FRUIT

EDU = 1 " " " " INCOME

= 2 " " " " EDUCATION

= 3 " " " " SOCIO-ECONOMIC STATUS

TEA = 0 NO ADJUSTMENT (OR MATCHING) FOR TEA

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
RANDOM-EFFECTS META-ANALYSES
Confounder Differences based on Unweighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	61.9518	(66)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0110	0.0211	N.S.	1.0110	0.9701	1.0537

SEPARATE RANDOM-EFFECTS META-ANALYSES BY REGION

NORTH AMERICA AND EUROPE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	35.3193	(36)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0188	0.0200	N.S.	0.9814	0.9437	1.0206

ASIA
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	23.7372	(29)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0549	0.0393	N.S.	1.0565	0.9782	1.1410

USA AND CANADA
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	19.4670	(20)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0313	0.0240	N.S.	0.9691	0.9246	1.0158

EUROPE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	14.6570	(15)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0096	0.0367	N.S.	1.0096	0.9395	1.0850

CHINA AND HONG KONG
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	14.7459	(17)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0407	0.0586	N.S.	1.0415	0.9286	1.1682

REST OF ASIA

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY REGION
Confounder Differences based on Unweighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	9.7461	(11)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0809	0.0382	(+)	1.0843	1.0061	1.1685

SEPARATE RANDOM-EFFECTS META-ANALYSES BY YEAR OF PUBLICATION

PUBLISHED IN 1980S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	20.6069	(24)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0755	0.0284	+	1.0784	1.0201	1.1400

PUBLISHED IN 1990S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	27.1304	(26)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0290	0.0330	N.S.	0.9714	0.9107	1.0363

PUBLISHED IN 2000S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	13.8624	(14)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0125	0.0404	N.S.	1.0126	0.9355	1.0960

SEPARATE RANDOM-EFFECTS META-ANALYSES BY NUMBER OF LUNG CANCER CASES

<100 CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	38.0428	(38)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0516	0.0336	N.S.	1.0529	0.9857	1.1247

100-199 CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	12.5163	(13)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0113	0.0273	N.S.	0.9888	0.9374	1.0430

200-399 CASES

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY NUMBER OF LUNG CANCER CASES
Confounder Differences based on Unweighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	7.9974	(8)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0884	0.0355	+	1.0925	1.0191	1.1711

400+ CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	4.0646	(4)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.1142	0.0601	N.S.	0.8920	0.7929	1.0036

SEPARATE RANDOM-EFFECTS META-ANALYSES BY DOSE-RESPONSE OR NOT

RESULTS FOR DOSE-RESPONSE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	21.7510	(22)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0388	0.0261	N.S.	1.0395	0.9877	1.0940

NO RESULTS FOR DOSE-RESPONSE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	39.4237	(43)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0084	0.0297	N.S.	0.9917	0.9356	1.0511

SEPARATE RANDOM-EFFECTS META-ANALYSES BY AGE ADJUSTMENT OR NOT

ADJUSTED (OR MATCHED) FOR AGE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	47.9801	(54)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0191	0.0201	N.S.	0.9811	0.9433	1.0205

NOT ADJUSTED (OR MATCHED) FOR AGE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	12.2295	(11)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.1688	0.0600	+	1.1839	1.0524	1.3318

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY STUDY TYPE
Confounder Differences based on Unweighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

CASE-CONTROL STUDIES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	53.1419	(56)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0100	0.0245	N.S.	1.0100	0.9627	1.0596

PROSPECTIVE STUDIES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	4.3328	(9)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0149	0.0368	N.S.	1.0150	0.9444	1.0909

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Weighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	RR	RRL	RRU	CA1	CA0	CO1	CO0	NCigs
Akiba	Japan	1.2521	0.5876	2.5154	22.9803	16.1244	66.6768	58.5780	1.0000
Akiba	Japan	1.3844	0.6332	3.0183	17.3431	16.1244	45.5107	58.5780	2.1350
Akiba	Japan	1.8180	0.4639	7.4310	3.5590	16.1244	7.1119	58.5780	3.8390
Boffetta 2	7 countries	0.9193	0.4472	1.7954	22.7678	24.4081	49.3217	48.6089	1.8175
Boffetta	7 countries	0.9432	0.7219	1.2430	161.3102	159.8555	306.3433	286.3355	1.3380
Boffetta	7 countries	0.5090	0.3002	0.8402	21.6909	159.8555	76.3271	286.3355	2.5880
Boffetta	7 countries	1.1101	0.6745	1.8326	29.3108	159.8555	47.2945	286.3355	4.3060
Brownson 1	USA	1.6330	0.3524	7.2159	3.3619	11.7639	5.5727	31.8438	2.3157
Brownson 2	USA	0.8155	0.6341	1.0069	182.7453	212.8397	548.8626	521.3277	2.3157
Buffler	USA	0.6020	0.2293	1.5953	21.4045	6.9378	113.3699	22.1208	2.3157
Butler	USA	1.9404	0.4485	8.4531	2.6313	5.5186	10777.1559	43858.5236	2.3157
Cardenas	USA	1.0403	0.4598	2.1396	8.4846	27.7004	10900.3459	37022.3902	1.0040
Cardenas	USA	1.0580	0.5794	2.0660	14.4570	27.7004	18262.9305	37022.3902	2.2670
Cardenas	USA	1.4951	0.6979	3.1943	8.7234	27.7004	7798.1436	37022.3902	4.3140
Chan	China	0.6070	0.3196	1.1457	22.6965	41.3570	50.3277	55.6655	1.8175
Choi	Korea	1.5650	0.8729	2.7885	45.5295	25.1251	82.9015	71.5967	1.8175
Correa	USA	1.7496	0.5248	5.7884	7.6800	5.1811	35.6602	42.0907	2.3157
De Waard	Netherlands	2.4680	0.7710	7.8865	17.0598	3.7341	112.1168	60.5647	1.8175
Du	China	0.6215	0.2895	1.3339	12.0933	24.8514	61.5368	78.5868	1.0000
Du	China	1.2098	0.6028	2.4279	18.8500	24.8514	49.2708	78.5868	2.6030
Enstrom	USA	0.8855	0.6153	1.2659	94.3638	42.5202	14340.0830	5721.6048	2.3157
Fontham	USA	1.1262	0.8895	1.4259	287.6932	162.4067	529.6106	336.7106	2.3157
Gao	China	1.1962	0.7812	1.8178	149.3323	44.7784	217.7827	78.1177	1.8175
Garfinkel 1	USA	1.2367	0.8344	1.8471	38.7551	65.2032	38365.7100	79824.0087	1.0040
Garfinkel 1	USA	1.0226	0.6995	1.4931	45.2382	65.2032	54155.6008	79824.0087	2.7520
Garfinkel 2	USA	0.8119	0.3832	1.7206	10.8459	41.8605	42.9472	134.5819	1.0040
Garfinkel 2	USA	0.9965	0.5786	1.7165	28.6966	41.8605	92.5813	134.5819	2.2670
Garfinkel 2	USA	1.7045	0.9319	3.1173	24.0735	41.8605	45.4082	134.5819	4.3140
Geng	China	1.3676	0.4635	4.0550	8.0445	11.9842	15.1547	30.8751	0.4850
Geng	China	1.8337	0.5186	6.4758	5.7253	11.9842	8.0440	30.8751	1.2730
Geng	China	2.4074	0.9130	6.3698	14.9077	11.9842	15.9537	30.8751	2.6030
Gorlova	USA	1.0494	0.5572	1.9771	56.6383	27.5079	58.7932	29.9642	2.3157
Hirayama	Japan	1.2956	0.8745	1.9282	98.5699	32.6549	43421.1840	18636.4922	0.8600
Hirayama	Japan	1.4038	0.8833	2.2359	39.0139	32.6549	15861.4250	18636.4922	2.6030
Hole	Scotland	1.2851	0.1152	14.5239	2.3107	0.9144	753.7820	383.3030	0.2080
Hole	Scotland	2.3711	0.1747	31.8679	1.4780	0.9144	261.3054	383.3030	2.4090
Humble 1	USA	1.4875	0.3286	7.0885	4.4345	6.0864	6.6468	13.5704	1.5820
Humble 1	USA	0.7973	0.0976	7.1656	1.4098	6.0864	3.9423	13.5704	3.3920
IARC: Kreuzer	Germany	0.7348	0.4509	1.2162	46.6918	44.0296	122.5265	84.9010	1.8175
Inoue	Japan	2.5340	0.2997	6.6161	6.6346	3.1457	11.8525	14.2407	1.0000
Inoue	Japan	2.9487	0.7691	15.1865	8.6488	3.1457	13.2780	14.2407	2.6030
Janerich	USA	0.5982	0.3588	0.9999	56.1972	63.3563	71.3995	48.1536	2.3157
Jee	Korea	1.9533	1.0670	3.8351	52.7559	11.4158	90421.9272	38219.7798	1.0000
Jee	Korea	1.4106	0.6467	3.1590	13.1237	11.4158	31148.0831	38219.7798	2.6030
Johnson	Canada	0.9840	0.4585	2.0911	25.1909	10.2782	271.2638	108.9051	1.8175
Kabat 1	USA	0.6740	0.1978	2.2535	10.4416	10.3286	12.9810	8.6543	2.3157
Kabat 2	USA	0.7069	0.3421	1.4692	14.7224	28.8263	48.3534	66.9296	1.6540
Kabat 2	USA	0.7649	0.3120	1.8807	8.2898	28.8263	25.1642	66.9296	3.6380
Kalandidi	Greece	1.5056	0.7715	2.9384	33.3956	26.2969	38.0751	45.1413	1.3380
Kalandidi	Greece	1.6790	0.7782	3.6225	21.0988	26.2969	21.5715	45.1413	2.9020
Kalandidi	Greece	1.4283	0.4851	4.2056	7.5274	26.2969	9.0466	45.1413	5.3350
Koo	China	2.2562	0.8705	5.8668	12.4274	21.7894	11.3309	44.8241	0.7080
Koo	China	1.6057	0.7055	3.6664	16.4024	21.7894	21.0138	44.8241	1.7670
Koo	China	1.0297	0.3587	2.9095	6.9163	21.7894	13.8174	44.8241	3.1830
Lagarde	Sweden	1.0243	0.7260	1.4503	66.6245	106.1266	239.4667	390.7100	1.8175
Lam T	China	2.1110	1.0868	4.0819	21.2736	75.7360	21.0781	158.4087	0.7080
Lam T	China	1.7073	1.0562	2.7539	45.1221	75.7360	55.2792	158.4087	1.7670

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Weighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	RR	RRL	RRU	CA1	CA0	CO1	CO0	NCigs
Lam T	China	1.7913	0.8631	3.7411	15.4116	75.7360	17.9954	158.4087	3.1830
Lam W	China	1.8603	0.9478	3.6645	29.3880	19.7472	52.4095	65.5133	1.8175
Layard	USA	0.5592	0.2037	1.5353	4.6218	22.2889	315.8944	851.8691	0.8190
Layard	USA	0.4956	0.1950	1.2593	5.6168	22.2889	433.1167	851.8691	2.6640
Lee	England	0.8297	0.2753	2.5074	13.2494	7.4523	29.1122	13.5864	1.8175
Lee C-H	Taiwan	1.7994	1.2380	2.6145	140.1614	59.1138	188.1217	142.7643	1.8175
Liu Q	China	0.6597	0.2061	2.1806	5.4265	11.8727	18.0705	26.0812	1.0000
Liu Q	China	2.4849	0.9016	7.1336	13.6749	11.8727	12.0889	26.0812	2.6030
Liu Z	China	0.6670	0.2445	1.8051	28.7033	6.3570	114.2689	16.8813	1.8175
Malats 7	Europe+Brazil	1.4022	0.7034	2.7835	32.6470	43.7959	21.9855	41.3550	1.8175
McGhee	China	1.2202	0.7934	1.8896	60.4124	64.3622	104.5597	135.9276	1.8175
Nishino	Japan	1.7295	0.6087	4.6743	8.2032	6.7286	19016.9502	26977.4906	1.8175
Ohno	Japan	0.9278	0.6133	1.4011	124.7549	50.6371	219.4829	82.6511	1.8175
Pershagen	Sweden	1.1076	0.6280	1.9942	31.3315	26.0694	125.4622	115.6218	1.8175
Rapiti	India	1.0820	0.4132	2.8533	10.0166	20.2781	15.5166	33.9889	1.8175
Rylander	Sweden	1.1761	0.4149	3.3406	5.3415	18.1506	30.6283	122.3997	1.8175
Schwartz	USA	0.9338	0.5931	1.4698	100.4456	62.4264	98.6016	57.2262	2.3157
Seow	Singapore	1.1780	0.8247	1.6828	75.7181	77.5576	261.2654	315.2603	1.8175
Shen	China	0.6261	0.1999	2.0172	11.1035	13.6371	12.4813	9.5975	0.4850
Shen	China	0.9516	0.2813	3.1582	11.5824	13.6371	8.5658	9.5975	1.2730
Shen	China	0.5725	0.2151	1.5320	24.7523	13.6371	30.4306	9.5975	2.6030
Shimizu	Japan	0.9867	0.5666	1.7159	43.6684	35.0218	79.5613	62.9573	1.8175
Sobue	Japan	1.0345	0.6956	1.5319	65.3657	53.7476	326.7333	277.9310	1.8175
Speize	USA	1.2646	0.2056	6.5340	18.5468	1.3797	191339.0630	18000.0000	2.3157
Stockwell	USA	1.4198	0.6711	2.8159	38.2485	17.6337	49.5733	32.4484	2.3157
Sun	China	1.0089	0.6709	1.5245	112.1250	76.8136	111.7183	77.2193	1.8175
Svensson	Sweden	1.2208	0.4423	3.3697	14.6219	6.3040	70.1635	36.9282	1.8175
Trichopoulos	Greece	0.5613	0.1208	2.6082	1.9972	24.3310	15.7953	108.0081	0.7080
Trichopoulos	Greece	2.4281	1.2232	4.8198	21.4820	24.3310	39.2740	108.0081	1.7670
Trichopoulos	Greece	3.8124	1.2553	11.5785	6.8617	24.3310	7.9897	108.0081	2.5880
Trichopoulos	Greece	1.7453	0.6470	4.7077	6.7639	24.3310	17.2039	108.0081	4.3060
Vineis	Western Europe	0.9838	0.5001	1.9503	19.5915	14.3729	27038.5563	19515.4318	1.8175
Wang L	China	1.0213	0.5925	1.6925	122.5358	24.9215	248.4654	51.6100	1.8175
Wang S	China	2.4409	1.2050	4.9636	64.2060	14.9057	57.8904	32.8045	1.8175
Wang T	China	0.3373	0.1047	1.1323	4.1494	40.3831	12.6371	41.4861	0.4850
Wang T	China	1.2253	0.6496	2.2994	39.7821	40.3831	33.3528	41.4861	1.2730
Wang T	China	1.1484	0.5822	2.2573	29.9846	40.3831	26.8231	41.4861	2.6030
Wu	USA	1.0891	0.4231	3.2121	14.7696	9.0408	28.0625	18.7082	2.3157
Wu-Williams	China	0.5850	0.4873	0.7741	206.6794	289.2018	393.6217	322.2337	1.8175
Zaridze	Russia	1.6264	1.0667	2.4719	61.0680	79.7559	86.2738	183.2597	0.7080
Zaridze	Russia	1.2636	0.7833	2.0481	36.7971	79.7559	66.9126	183.2597	2.2930
Zatloukal	Czech Republic	0.2626	0.0630	1.0876	2.0635	37.2154	72.4790	343.2232	1.8175
Zheng	China	2.3810	0.9372	6.0664	48.0612	5.7505	139.6019	39.7712	1.8175
Zhong	China	1.3436	0.8521	2.1403	79.1249	59.5256	77.2139	78.0470	0.7080
Zhong	China	0.8122	0.5234	1.3070	65.0742	59.5256	105.0480	78.0470	1.7670
Zhong	China	1.1637	0.5474	2.2973	18.1014	59.5256	20.3948	78.0470	3.1830

NOTES: RR,RL,RU RELATIVE RISKS AND LOWER AND UPPER CONFIDENCE INTERVALS

CA1,CA0 NUMBERS (OR PSEUDO-NUMBERS) OF EXPOSED AND UNEXPOSED CASES

CO1,CO0 NUMBERS (OR PSEUDO-NUMBERS) OF EXPOSED AND UNEXPOSED CONTROLS

NCIGS MIDPOINT OF NUMBER OF CIGARETTES SMOKED BY HUSBAND (IN UNITS OF 10)

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

BASIC DATA USED IN META-ANALYSES

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION

Confounder Differences based on Weighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	Beta	St.Err. Beta	Z	Weight	ETSse	FRU	VEG	FAT	EDU	TEA
Akiba	Japan	0.1491	0.1468	1.0157	46.4032	0.9896	0	0	0	0	0
Boffetta 2	7 countries	-0.0463	0.1951	-0.2373	26.2715	0.9130	0	0	0	0	0
Boffetta	7 countries	-0.0359	0.0532	-0.6748	353.3269	1.1768	0	0	0	0	0
Brownson 1	USA	0.2118	0.3326	0.6368	9.0397	0.8345	0	0	0	1	0
Brownson 2	USA	-0.0881	0.0509	-1.7308	385.9797	1.1579	0	0	0	0	0
Buffler	USA	-0.2192	0.2137	-1.0257	21.8973	0.8581	0	0	0	0	0
Butler	USA	0.2863	0.3235	0.8850	9.5555	0.9215	0	0	0	0	0
Cardenas	USA	0.0793	0.0844	0.9396	140.3832	1.4556	0	1	1	2	0
Chan	China	-0.2747	0.1792	-1.5329	31.1404	0.9109	0	0	0	0	0
Choi	Korea	0.2464	0.1630	1.5117	37.6378	0.9091	0	0	0	0	0
Correa	USA	0.2416	0.2645	0.9134	14.2938	1.1582	0	0	0	0	0
De Waard	Netherlands	0.4971	0.3264	1.5230	9.3864	0.8696	0	0	0	0	0
Du	China	0.0804	0.1365	0.5890	53.6704	1.0879	0	0	0	0	0
Enstrom	USA	-0.0525	0.0795	-0.6604	158.2216	1.0456	1	0	0	2	0
Fontham	USA	0.0513	0.0520	0.9865	369.8225	1.1293	1	1	0	2	0
Gao	China	0.0986	0.1185	0.8321	71.2137	0.8022	0	0	0	2	0
Garfinkel 1	USA	0.0045	0.0702	0.0641	202.9204	1.1963	0	0	0	0	0
Garfinkel 2	USA	0.1127	0.0689	1.6357	210.6500	1.5345	0	0	0	0	0
Geng	China	0.3258	0.1828	1.7823	29.9259	1.0478	0	0	0	0	0
Gorlova	USA	0.0208	0.1395	0.1491	51.3868	1.1005	0	0	0	1	0
Hirayama	Japan	0.1045	0.0864	1.2095	133.9592	0.8998	0	0	0	0	0
Hole	Scotland	0.3071	0.4358	0.7047	5.2653	1.0530	0	0	0	0	0
Humble 1	USA	0.0063	0.3010	0.0209	11.0374	1.2885	0	0	0	0	0
IARC: Kreuzer	Germany	-0.1695	0.1393	-1.2168	51.5345	0.8957	0	0	0	0	0
Inoue	Japan	0.3357	0.2749	1.2212	13.2328	1.1196	0	0	0	0	0
Janerich	USA	-0.2219	0.1129	-1.9655	78.4535	1.1397	0	0	0	0	0
Jee	Korea	0.0516	0.1508	0.3422	43.9741	0.9132	0	1	0	3	0
Johnson	Canada	-0.0089	0.2130	-0.0418	22.0415	0.8225	1	1	0	2	0
Kabat 1	USA	-0.1704	0.2680	-0.6358	13.9229	1.1579	0	0	0	0	0
Kabat 2	USA	-0.0944	0.1213	-0.7782	67.9639	1.3646	0	0	0	2	0
Kalandidi	Greece	0.0953	0.0916	1.0404	119.1816	1.5584	0	0	0	0	0
Koo	China	0.0478	0.1533	0.3118	42.5516	1.1884	0	0	0	2	0
Lagarde	Sweden	0.0132	0.0971	0.1359	106.0624	0.8828	0	0	0	3	0
Lam T	China	0.2243	0.0984	2.2795	103.2785	1.0055	0	0	0	0	0
Lam W	China	0.3415	0.1898	1.7993	27.7592	0.9063	0	0	0	0	0
Layard	USA	-0.2760	0.1778	-1.5523	31.6327	1.1466	0	0	0	0	0
Lee	England	-0.1027	0.3101	-0.3312	10.3991	0.8544	0	0	0	0	0
Lee C-H	Taiwan	0.3232	0.1049	3.0810	90.8760	0.9015	0	0	0	2	0
Liu Q	China	0.3645	0.2024	1.8009	24.4106	1.0561	0	0	0	2	0
Liu Z	China	-0.2228	0.2806	-0.7940	12.7006	0.6106	0	0	0	0	0
Malats	7 Europe+Brazil	0.1860	0.1931	0.9632	26.8186	0.8717	0	0	0	0	0
McGhee	China	0.1095	0.1218	0.8990	67.4071	0.9025	0	0	0	2	0
Nishino	Japan	0.3014	0.2861	1.0535	12.2170	0.8950	1	1	1	0	0
Ohno	Japan	-0.0413	0.1160	-0.3560	74.3163	0.8114	0	0	0	0	0
Pershagen	Sweden	0.0562	0.1622	0.3465	38.0100	0.9097	0	0	0	0	0
Rapiti	India	0.0434	0.2712	0.1600	13.5963	0.8504	0	0	0	0	0
Rylander	Sweden	0.0892	0.2928	0.3046	11.6643	0.7290	0	0	0	0	0
Schwartz	USA	-0.0296	0.1000	-0.2960	100.0000	1.1194	0	0	0	0	0
Seow	Singapore	0.0902	0.1001	0.9011	99.8003	0.9054	0	0	0	0	0
Shen	China	-0.1607	0.1676	-0.9588	35.6002	1.0903	0	0	0	0	0
Shimizu	Japan	-0.0074	0.1555	-0.0476	41.3561	0.9053	0	0	0	0	0
Sobue	Japan	0.0187	0.1108	0.1688	81.4555	0.9064	0	0	0	2	0
Speize	USA	0.1014	0.3811	0.2661	6.8853	0.6375	0	0	0	0	0
Stockwell	USA	0.1514	0.1580	0.9582	40.0577	1.1381	0	0	0	2	0
Sun	China	0.0049	0.1152	0.0425	75.3520	0.8954	0	0	0	2	0
Svensson	Sweden	0.1098	0.2850	0.3853	12.3115	0.8673	0	0	0	0	0
Trichopoulos	Greece	0.2442	0.1071	2.2801	87.1808	1.3389	0	0	0	0	0

ALL STUDIES

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Confounder Differences based on Weighted Regression

Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Study title	Country	Beta	St.Err. Beta	Z	Weight	ETSse	FRU	VEG	FAT	EDU	TEA
Vineis	Western Europe	-0.0090	0.1910	-0.0471	27.4115	0.8968	1	1	1	2	0
Wang L	China	0.0116	0.1473	0.0788	46.0887	0.6870	0	0	0	1	0
Wang S	China	0.4910	0.1987	2.4711	25.3282	0.8780	0	0	0	0	0
Wang T	China	0.0970	0.1308	0.7416	58.4500	1.0379	0	0	0	0	0
Wu	USA	0.0369	0.2233	0.1652	20.0550	1.1449	0	0	0	0	0
Wu-Williams	China	-0.2949	0.0650	-4.5369	236.6864	0.9047	0	0	0	2	0
Zaridze	Russia	0.1046	0.1069	0.9785	87.5074	0.8744	0	0	0	2	0
Zatloukal	Czech Republic	-0.7358	0.3999	-1.8400	6.2531	0.6900	0	0	0	2	0
Zheng	China	0.4773	0.2621	1.8211	14.5568	0.7567	0	0	0	0	0
Zhong	China	-0.0646	0.0971	-0.6653	106.0624	0.9343	1	0	0	1	0

NOTES: BETA SLOPE OF LOG RR ON NCIGS

SE BETA STANDARD ERROR OF BETA

Z RATIO OF BETA TO ITS STANDARD ERROR (APPROXIMATE NORMAL STATISTIC)

WEIGHT INVERSE OF THE VARIANCE OF BETA

ETSse STANDARD ERROR OF ETS EXPOSURE OVER THE CONTROL GROUPS

FRU = 1 DOSE-RESPONSE ANALYSIS ADJUSTED FOR FRUIT

VEG = 1 " " " " VEGETABLES

FAT = 1 " " " " FRUIT

EDU = 1 " " " " INCOME

= 2 " " " " EDUCATION

= 3 " " " " SOCIO-ECONOMIC STATUS

TEA = 0 NO ADJUSTMENT (OR MATCHING) FOR TEA

ALL STUDIES

EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY

GENG DATA REPLACED WITH PNL ESTIMATES

WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
RANDOM-EFFECTS META-ANALYSES
Confounder Differences based on Weighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	61.9865	(66)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0387	0.0206	(+)	1.0394	0.9983	1.0822

SEPARATE RANDOM-EFFECTS META-ANALYSES BY REGION

NORTH AMERICA AND EUROPE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	35.9205	(36)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0059	0.0186	N.S.	1.0059	0.9699	1.0432

ASIA
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	23.6538	(29)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0851	0.0385	+	1.0888	1.0096	1.1742

USA AND CANADA
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	19.3584	(20)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0070	0.0226	N.S.	0.9931	0.9501	1.0379

EUROPE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	14.7794	(15)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0363	0.0347	N.S.	1.0369	0.9687	1.1100

CHINA AND HONG KONG
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)				
Model 1	14.7583	(17)				
	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0708	0.0575	N.S.	1.0734	0.9589	1.2014

REST OF ASIA

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY REGION
Confounder Differences based on Weighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	9.2218	(11)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.1128	0.0381	+	1.1194	1.0389	1.2062

SEPARATE RANDOM-EFFECTS META-ANALYSES BY YEAR OF PUBLICATION

PUBLISHED IN 1980S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	19.9280	(24)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.1036	0.0283	++	1.1091	1.0492	1.1725

PUBLISHED IN 1990S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	27.1516	(26)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0012	0.0320	N.S.	0.9988	0.9381	1.0634

PUBLISHED IN 2000S
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	13.8006	(14)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0397	0.0400	N.S.	1.0405	0.9621	1.1254

SEPARATE RANDOM-EFFECTS META-ANALYSES BY NUMBER OF LUNG CANCER CASES

<100 CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	38.0144	(38)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0793	0.0328	+	1.0825	1.0150	1.1545

100-199 CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	12.3891	(13)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0170	0.0272	N.S.	1.0172	0.9643	1.0729

200-399 CASES

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY NUMBER OF LUNG CANCER CASES
Confounder Differences based on Weighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	7.9442	(8)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.1153	0.0352	+	1.1222	1.0474	1.2023

400+ CASES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	4.0518	(4)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	-0.0839	0.0573	N.S.	0.9196	0.8218	1.0290

SEPARATE RANDOM-EFFECTS META-ANALYSES BY DOSE-RESPONSE OR NOT

RESULTS FOR DOSE-RESPONSE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	21.7945	(22)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0654	0.0253	+	1.0676	1.0160	1.1218

NO RESULTS FOR DOSE-RESPONSE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	39.4466	(43)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0195	0.0290	N.S.	1.0197	0.9633	1.0794

SEPARATE RANDOM-EFFECTS META-ANALYSES BY AGE ADJUSTMENT OR NOT

ADJUSTED (OR MATCHED) FOR AGE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	48.4986	(54)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0086	0.0194	N.S.	1.0087	0.9710	1.0478

NOT ADJUSTED (OR MATCHED) FOR AGE
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	12.1556	(11)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.1960	0.0589	++	1.2165	1.0839	1.3654

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1

APPENDIX 15

ETS and Lung Cancer - Meta-Analysis of Dose Response Exposure (or nearest equivalent), 08-SEP-06

Female (Husband's smoking) - Number cigarettes smoked

CORRECTED FOR ALL CONFOUNDERS (FRUIT, VEG, FAT, EDUCATION) AND MISCLASSIFICATION
SEPARATE RANDOM-EFFECTS META-ANALYSES BY STUDY TYPE
Confounder Differences based on Weighted Regression
Misclassification: Additive, Concordance: 3.0, MissP: US:0.025, EU:0.025, ASIA:0.1

CASE-CONTROL STUDIES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	53.1173	(56)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0388	0.0239	N.S.	1.0396	0.9919	1.0895

PROSPECTIVE STUDIES
Dose Response (Beta)
WEIGHTED on Beta Weights

	Deviance	(DF)
Model 1	4.2697	(9)

	Estimate	S.E.	P	RR	95%CIl	95%CIu
Constant	0.0385	0.0367	N.S.	1.0392	0.9670	1.1168

ALL STUDIES
EXPOSURE GROUPS BASED ON ISSNZ, MODIFIED FOR USA, EXPOSURE IN 10 CIGS/DAY
GENG DATA REPLACED WITH PNL ESTIMATES
WEIGHT ON (1/se*se), Variance set at 1