

Philosophy 692S  
**Bioethics: Advanced Research Ethics**  
Fall 2013

Monday 3:05-5:35  
Seeley Mudd 108

Trent Center for Bioethics

**Schedule**

Monday, August 26	Introduction (GS & RM)
Mon, September 2	Conflict of interest I (RM)
Mon, Sept 9	Conflict of interest II (RM)
Mon, Sept 16	case: incentives, guinea pigging, (RM)
Mon, Sept 23	Clinical equipoise I: orthodoxy & critique (GS)
Mon, Sept 30	Clinical equipoise II: defence (GS)
Mon, October 7	case: Abigail Alliance (Peppercorn)
Mon, Oct 14	<i>Fall break. No class.</i>
Mon, Oct 21	Informed consent I: conventional wisdom (GS)
Mon, Oct 28	Therapeutic misconception (Weinfurt) <b>Graduate student topic proposals due</b>
Mon, November 4	Informed consent II: understanding required? (GS)
Mon, Nov 11	Learning health systems (Califf)
Mon, Nov 18	Adaptive design; Bayesianism (Danny Benjamin)
Mon, Nov 25	case: SUPPORT (RM)

## Readings

- Aug 26      Nothing.
- Sept 2      S. Heres et al., "Why Olanzapine beats risperidone, risperidone beats quetiapine, and quetiapine beats olanzapine: An Exploratory analysis of head-to-head comparison studies of second-generation antipsychotics," *Am J Psychiatry* 163 (2006): 185-194;  
B. Als-Nielsen et al., "Association of funding and conclusions in randomized drug trials: A Reflection of treatment effect or adverse events?," *JAMA* 290 (2003): 921-928;  
EH Turner et al., "Selective publication of antidepressant trials and its influence on apparent efficacy," *NEJM* 358 (2008): 252-60;  
Multiple authors: AAMC. "The Scientific Basis of influence and reciprocity: A Symposium," *AAMC Publications*, (2007) Read pp 3-28. 29-34 optional.
- Sept 9      KP Weinfurt et al., "Disclosure of financial relationships to participants in clinical research," *NEJM* 331 (2009): 916-921;  
AS Kesselheim et al., "A randomized study of how physicians interpret research funding disclosures," *NEJM* 367 (2012): 1119-27;  
H Bauchner, PB Fontanarosa, "Restoring confidence in the pharmaceutical industry," *JAMA* 309 (2013): 607-609
- Sept 16     C. Elliott, "Guinea Pigging," *The New Yorker* (2008, January 7);  
C. Grady, "Payment of clinical research subjects," *J Clin Invest* 115 (2005): 1681-1687  
E. Largent et al., "Money, coercion, and undue inducement: Attitudes about payments to research participants," *IRB* 34 (2012): 1-8  
JP Bentley, PG Thacker, "The influence of risk and monetary payment on the research participation decision making process," *J Med Ethics* 30 (2004): 293-98
- Sept 23     B Freedman, "Equipoise and the ethics of clinical research," *N Engl J Med* 317 (1987): 141-5;  
B Freedman, "Placebo-Controlled Trials and the Logic of Clinical Purpose," *IRB* 12 (6) (1990): 1-6  
FG Miller, H Brody, "A critique of clinical equipoise: therapeutic misconception in the ethics of clinical trials," *Hastings Center Report* 33 (3) (2003): 19-28;  
FG Miller, S Joffe, "Equipoise and the Dilemma of Randomized Clinical Trials," *N Engl J Med* 364 (2011): 476-80
- Sept 30     Hawkins and Emanuel, *Exploitation and developing countries* (Princeton, 2008): 36-44.

- Gifford, "Community equipoise and the ethics of randomized clinical trials," *Bioethics* 9 (1995): 127-48;
- Gifford, "Freedman's 'Clinical Equipoise' and 'Sliding-Scale All-Dimensions-Considered Equipoise'," *Journal of Medicine and Philosophy* 25 (2000): 399-426.
- Oct 7      Peppercorn et alia, "The dilemma of data-safety monitoring: provision of significant new data to research participants," *Lancet* 371 (2008): 527-29;
- Peppercorn et alia, "Self-reported practices and attitudes of US oncologists regarding off-protocol therapy," *Journal of Clinical Oncology* 26 (36) (2008): 5994-6000;
- Peppercorn et alia, "Ethics of mandatory research biopsy for correlative end points within clinical trials in oncology," *Journal of Clinical Oncology* 28 (15) (2010): 2635-40.
- Oct 21      Faden and Beauchamp, *History and Theory of Informed Consent* (Oxford, 1986): chh. 8-9.
- Berg et alia, *Informed Consent: Legal Theory and Clinical Practice*, 2 ed. (Oxford, 2001), ch. 3
- Oct 28      Appelbaum et alia, "False Hopes and Best Data: Consent to Research and the Therapeutic Misconception," *Hastings Center report* 17(2) (1987): 20-24;
- Hornig & Grady, "Misunderstanding in clinical research: Distinguishing therapeutic misconception, therapeutic misestimation, & therapeutic optimism," *IRB* (2003): 11-16;
- Kim, S., Schrock, L., Wilson, R., Frank, S., Holloway, R., Kiebertz, K., & de Vries, R, "An approach to evaluating the therapeutic misconception," *IRB* (2009);
- Lidz & Appelbaum, "The therapeutic misconception: problems and solutions," *Medical Care* 40(9) (2002).
- Nov 4      Sreenivasan, "Does informed consent to research require comprehension?," *The Lancet* 362 (2003): 2016-18.
- A. Meisel, "The 'exceptions' to the informed consent doctrine," *Wisconsin Law Review* (1979): 454
- Nov 11      Faden et alia, "An Ethics Framework for a Learning Health System," *Hastings Center Report* 43(1) (2013): S16-S27;
- Kass et alia, "The Research-Treatment Distinction," *Hastings Center Report* 43(1) (2013): S4-S15;
- IOM, *Better Care at Lower Cost* (summary).

- Nov 18 N. Silver, *The Signal and the noise* (Penguin, 2012), ch. 8;  
TR Fleming, JH Powers, "Issues in noninferiority trials: the evidence in  
community-acquired pneumonia," *Clin Infect Dis.* (2008) Dec 1;47  
Suppl 3:S108-20. doi: 10.1086/591390  
Benjamin et alia, "Safety and transparency of pediatric drug trials," *Arch Pediatr  
Adolesc Med.* 163 (2009): 1080–1086
- Nov 25 D. Magnus, A. Caplan, "Risk, Consent, and SUPPORT," *N Engl J Med* 368  
(2013): 1864-65.

### **Assignments**

Everyone will be required to write a weekly 600 word essay on the assigned topic. Essays will be due at noon in the Sakai dropbox on Mondays (ie, class day). Topics will be posted in Sakai the week before. Everyone has **two** free passes on the short essays, i.e. you may elect not to submit an essay on two weeks of your choosing.

Graduate students will also be required to write a 3000 word essay at the end of term. Your 1 page topic proposal is due by e-mail on Monday, October 28. The essay itself is due Monday, December 9.

### **Grading**

The essays will be graded by Professors McKinney and Sreenivasan.

On the short essays, we will count your highest 8 grades (out of the ten essays submitted).

For graduate students, these 8 grades will count for 60 percent of your final grade; your longer essay will count for 20 percent; and participation in class will also count for 20 percent.

For undergraduates, the 8 short essays grades will count for 75 percent of your final grade and participation will count for 25 percent. There is no longer essay required. However, any undergraduate may elect to follow the graduate pattern.

### **Office hours**

The primary instructors are available by appointment.

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Gopal Sreenivasan                [gopal.sreenivasan@duke.edu](mailto:gopal.sreenivasan@duke.edu)

16 September, 2013