

An Overview of Hybrid Effectiveness-Implementation Designs

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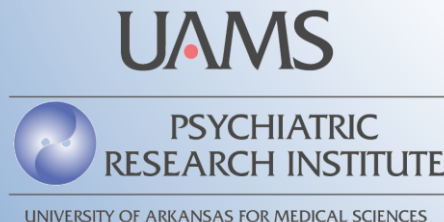
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Goals for the session

- Discuss the concept of “hybrid designs” which combine elements of clinical/preventive effectiveness and implementation research
 - Type 1: Explore *Implementability* of an intervention while we are testing its effectiveness (towards real world implementation strategies)
 - Type 2: Test implementation strategies *during* effectiveness trials (simultaneous look at both)
 - Type 3: Test implementation strategies while also documenting clinical/prevention intervention outcomes (evaluating them as they relate to uptake and fidelity)
- Review trends in use of designs; some examples
- Present newer thinking on specification, measurement, reporting

The next few slides cover material from this paper:



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Med Care. Author manuscript; available in PMC 2013 August 01.

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**Effectiveness-implementation Hybrid Designs:
Combining Elements of Clinical Effectiveness and Implementation Research to Enhance
Public Health Impact**

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Some of the later slides are reflected in this paper



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An introduction to effectiveness-implementation hybrid designs

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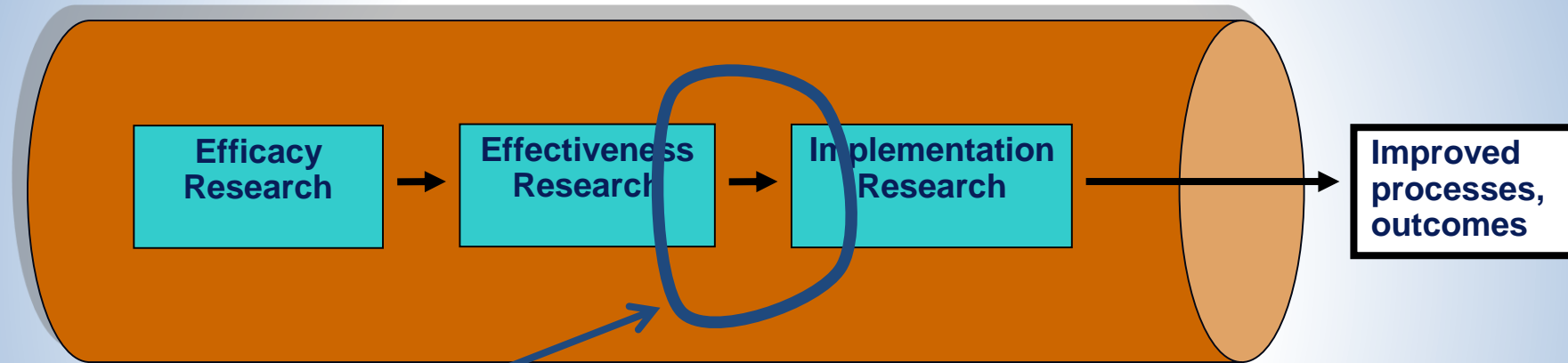
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Why Hybrid Designs?

- The speed of moving research findings into routine adoption could be improved by considering *hybrid designs* that combine elements of effectiveness and implementation research
 - Or, combine research questions in both areas
- Don't wait for “perfect” effectiveness data before moving to implementation research
- We can “backfill” effectiveness data while we test implementation strategies
- How do clinical outcomes relate to levels of adoption and fidelity?
 - How will we know this without data from “both sides”?

Effectiveness-Implementation hybrid designs

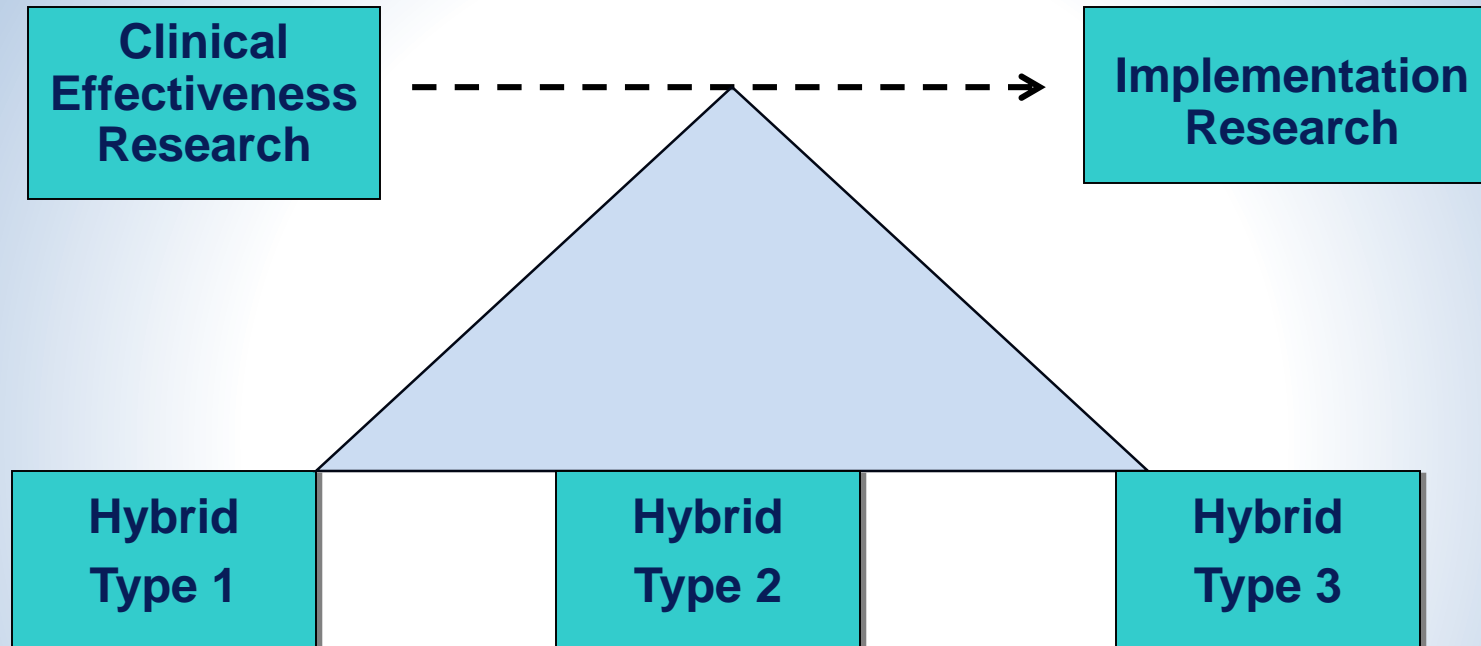


Spatially speaking, hybrids “fit” in here...

Couple of intro thoughts about hybrids

- Researchers were doing “hybrid designs” well before we began speaking and writing about them as such
- The paper in 2012 tried to bring some attention to the issues surrounding such combinations, along with some direction, examples, and recommendations
- Original paper focused on trials but these hybrid concepts can and are being used in other designs
 - *It's more about combining research questions*

Types of Hybrids



Hybrid Type 1: test the **thing**, observe/gather information on **doing the thing**

Hybrid Type 2: test **thing**, test/study **do the thing**

Hybrid Type 3: test **do the thing**, observe/gather information on the **thing**

Hybrid Type 1 Designs

Definition:

- Test clinical intervention and explore implementation-related factors (80%/20%?)

Description:

- Conventional effectiveness study “plus”:
 - Describe implementation experience (worked/didn’t; barriers/facilitators)
 - How might the intervention need to be adapted going forward?
 - What is needed to support people/places to do THE THING in the real world?

Indications (circa 2012):

- Clinical effectiveness evidence remains limited, so intensive focus on implementation might be premature...BUT
- Effectiveness study conditions offer ideal opportunity to explore implementation issues, plan implementation strategies for next stage

Remember...

- All effectiveness trials use “implementation strategies” to support the delivery of the intervention; we just usually don’t call them that...
- They are normally resource-intensive
 - Paying clinics, paying interventionists, paying for care, frequent fidelity checks and intervening when it goes south...
- We “know” that some/many the strategies used in effectiveness trials are not feasible for supporting wide-spread adoption
- BUT, we can learn from the use of those strategies during the trial!

More Design Considerations: Type 1

- The original definition of a type 1 emphasized secondary aims/questions and exploratory data collection and analysis preparatory to a greater focus on implementation activity
 - Review indicates that this is the common model of type 1
- However, some type 1 studies are doing more intense focus on “implementability” in developing/adapting intervention before effectiveness trial
 - i.e., “(re-)design for dissemination/implementation” step first
- What if you have a small number of sites?
 - Expand data collection to *naïve* sites (clinics not yet doing the thing)

Example of Type 1: CALM study

- **Curran et al., 2012, *Implementation Science***
- **Large effectiveness trial of anxiety intervention in primary care**
 - 4 cities, 17 clinics, 1004 patients
 - Care managers using software tool with patients to navigate Tx manual
 - Care managers were local nurses/social workers already working in the clinics
 - Intervention was designed with “future implementation in mind”
- **Qualitative process evaluation alongside trial**
 - 47 interviews with providers, nurses, front office, and anxiety care managers
 - Most interviews done on the phone
 - Interview guide informed by an implementation framework (PARIHS)
 - (these days, that link needs to be very explicit...)

What did we learn?

- Lots of stuff...
- But, I'll share one important piece of data that illustrates the value of this kind of evaluation
 - Many of the providers in the participating clinics DID NOT refer a lot of patients for the trial. Some referred NOBODY.
 - Those who referred a lot were *already* interested in MH
 - Those who didn't were not persuaded during the site trainings that this was a good enough idea to actually take part
 - So, "uptake" and "reach" were not great in the trial, even though the researchers tried to get all providers to refer
 - So, key barrier to future implementation was provider buy-in and engagement. "Standard" strategies to entice them didn't work.
 - We would have learned this about this barrier about 2+ years later if we had done this sequentially.

More Type 1 examples

Contemporary Clinical Trials 43 (2015) 260–278

Contents lists available at ScienceDirect

 **Contemporary Clinical Trials**

journal homepage: www.elsevier.com/locate/conclintrial



Research aimed at improving both mood and weight (RAINBOW) in primary care: A type 1 hybrid design randomized controlled trial[☆]

Jun Ma^{a,b,*}, Veronica Yank^b, Nan Lv^a, Jeremy D. Goldhaber-Fiebert^c, Megan A. Lewis^d, M. Kaye Kramer^e, Mark B. Snowden^f, Lisa G. Rosas^{a,b}, Lan Xiao^a, Andrea C. Blonstein^a



Lane-Fall *et al. BMC Surgery* 2014, **14**:96
<http://www.biomedcentral.com/1471-2482/14/96>



STUDY PROTOCOL

Open Access

Handoffs and transitions in critical care (HATRICC): protocol for a mixed methods study of operating room to intensive care unit handoffs

Meghan B Lane-Fall^{1,2,3*}, Rinad S Beidas^{3,4}, Jose L Pascual^{5,6}, Meredith L Collard¹, Hannah G Peifer⁷, Tyler J Chavez⁸, Mark E Barry², Jacob T Gutsche¹, Scott D Halpern^{3,9,10}, Lee A Fleisher^{1,3,10} and Frances K Barg^{11,12}

Hagedorn *et al. Addiction Science & Clinical Practice* 2014, **9**:12
<http://www.ascpjournal.org/content/9/1/12>



RESEARCH

Open Access

An implementation-focused process evaluation of an incentive intervention effectiveness trial in substance use disorders clinics at two Veterans Health Administration medical centers

Hildi J Hagedorn^{1,2,3*}, Cheryl B Stetler⁴, Ann Bangerter², Siamak Noorbaloochi^{2,3}, Maxine L Stitzer⁵ and Daniel Kivlahan^{6,7}

Bramoweth *et al. Trials* (2018) 19:73
DOI 10.1186/s13063-017-2437-y

Trials

STUDY PROTOCOL

Open Access

A hybrid type I trial to increase Veterans' access to insomnia care: study protocol for a randomized controlled trial

Adam D. Bramoweth^{1,2*}, Anne Germain³, Ada O. Youk^{1,4}, Keri L. Rodriguez^{1,5} and Matthew J. Chinman^{1,2,6}



Hybrid Type 2 Designs

Definition:

- Test clinical intervention and test/study implementation strategy (50/50? 60/40? 72/28?)

Description:

- Dual-focus study:
 - Clinical Effectiveness trial within either:
 - Implementation trial of 2+ strategies
 - Pilot (non-randomized) study of single implementation strategy

Indications (circa 2012):

- Clinical effectiveness data available, though perhaps not for context/population of interest for this trial
- Data on barriers and facilitators to implementation available
- Implementation momentum in terms of system/policy demands?

More Design Considerations: Type 2

- The original definition of a type 2 described possibilities of dual focused, dual randomized designs & randomized effectiveness trials nested in pilots of an implementation strategy
 - Majority of currently published Type 2s are the latter
 - Some dual randomized designs (see example soon)
- When looking at the aims or hypotheses of existing studies, most have primary aim on *intervention outcomes*

More Design Considerations: Type 2

- Important to have an explicitly described implementation strategy that is thought to be plausible in the real world
 - Clear distinction from type 1
- Explicit measurement of adoption, fidelity...
 - Always happens in type 2
- Important to be clear about intervention components versus implementation strategy components
 - Existing papers sometimes not clear here
 - This isn't always easy to decide or describe
 - E.g., *delivery format...*
 - *Is delivering the intervention over the telephone an intervention component or an implementation strategy?*

Still More Design Considerations: Type 2

- **What if the implementation strategy leads to poor adoption and poor fidelity?**
 - Effectiveness trial gets compromised
- **What to do about this?**
 - Use implementation strategies with relevant evidence base
 - Build in adoption/fidelity benchmarks
 - Build in measurement and plans to address poor adoption and/or fidelity
 - Build in time to deal with this possibility
 - Anyone getting queasy over this?? Understandable....

Example 1: Cully et al., 2012, 2014+

- **Clinical trial of brief cognitive behavioral therapy in treating depression and anxiety; 1 “pilot” implementation strategy**
 - Patient randomization only; Pilot study of implementation strategy (online training, audit and feedback, facilitation) in 2 large VAMCs
 - Intent-to-treat analysis of clinical outcomes (N=320)
 - Feasibility, acceptability, and “preliminary effectiveness” data collected on implementation strategy
 - Measured knowledge acquisition, fidelity to model
 - Qualitative data on implementability, time spent, etc.
 - Measured sustainability of provision of brief CBT after trial
 - Preparatory to implementation trial of strategy

Example 2: Garner et al., 2017

- **Aim 1**: effectiveness of a motivational interviewing-based brief intervention (MIBI) for substance use as an adjunct to usual care (referral) within AIDS service organizations (ASOs)
- **Aim 2**: effectiveness of implementation and sustainment facilitation (ISF) as an adjunct to the Addiction Technology Transfer Center (ATTC) model for training staff in MI
 - Patients randomized within ASOs (N=1872)
 - SUD outcomes
 - ASOs randomized to ACCT or ACCT+ISF (N=39)
 - Proctor et al (2011) measures (*pretty much all of them...!*)

More Type 2 examples

TBM

ORIGINAL RESEARCH



Evaluating the effectiveness of physician counseling to promote physical activity in Mexico: an effectiveness-implementation hybrid study

Karla I. Galaviz,¹ Paul A. Estabrooks,² Edna Jauregui Ulloa,³ Rebecca E. Lee,⁴ Ian Janssen,⁵ Juan López y Taylor,⁶ Luis Ortiz-Hernández,⁷ Lucie Lévesque⁸

Rich et al. *BMC Public Health* (2018) 18:29
DOI 10.1186/s12889-017-4584-1

BMC Public Health

STUDY PROTOCOL

Open Access



Implementation-effectiveness trial of an ecological intervention for physical activity in ethnically diverse low income senior centers

Porchia Rich^{1*}, Gregory A. Aarons², Michelle Takemoto¹, Veronica Cardenas³, Katie Crist¹, Khalisa Bolling¹, Brittany Lewars¹, Cynthia Castro Sweet⁴, Loki Natarajan^{1,3}, Yuyan Shi¹, Kelsie M. Full¹, Eileen Johnson¹, Dori E. Rosenberg⁵, Melicia Whitt-Glover⁶, Bess Marcus¹ and Jacqueline Kerr^{1,3}

Open Access

Protocol

BMJ Open Effectiveness of implementing a best practice primary healthcare model for low back pain (BetterBack) compared with current routine care in the Swedish context: an internal pilot study informed protocol for an effectiveness-implementation hybrid type 2 trial

Allan Abbott,¹ Karin Schröder,¹ Paul Enthoven,¹ Per Nilsen,² Birgitta Öberg¹

Rogers et al. *Addiction Science & Clinical Practice* 2013, **8**:7
<http://www.ascjournal.org/content/8/1/7>



STUDY PROTOCOL

Open Access

Telephone care coordination for smokers in VA mental health clinics: protocol for a hybrid type-2 effectiveness-implementation trial

Erin Rogers^{1,2*}, Saida Fernandez⁸, Colleen Gillespie², David Smelson^{4,5}, Hildi J Hagedorn^{6,7}, Brian Elbel³, David Kalman^{4,5}, Alfredo Axtmayer¹, Karishma Kurowski^{1,2} and Scott E Sherman^{1,2}

Hybrid Type 3 Designs

Definition:

- test implementation strategy, observe/gather information on clinical intervention and outcomes

Description:

- Largely focused on trial of implementation strategies
- Randomization usually at level of provider, clinic, or system
- Clinical outcomes are “secondary”

Indications (circa 2012):

- We sometimes proceed with implementation studies without completing a “full portfolio” of effectiveness studies (e.g. mandates; VA anyone?)
 - Strong momentum in a system, e.g., “We are rolling this out!”
- Interested in exploring how clinical effectiveness might vary by level/quality of implementation?

More Design Considerations: Type 3

- How much power you got?
- Important to use outcomes framework
 - RE-AIM
 - Proctor et al., 2011
- What's your evidence for implementation strategies selected?
- What about *mechanisms of action* of the strategies?
- What about cost of the strategies?
 - Mechanisms and Cost will likely become essential parts of type 3 studies
- Clinical outcomes data collection
 - Do you really need them? What interventions might we NOT need to do a hybrid 3 study for?
 - Measures available in existing data?
 - Primary data collection? (*Mental Health outcomes not routinely available...*)
 - Sub-sample?

Smelson et al., 2015

- **Mission-Vet is an evidence-based treatment for co-occurring SUD and MH disorders among homeless Veterans**
- **Compare “implementation as usual” of Mission-Vet to IAU plus Getting To Outcomes (GTO)**
 - IAU = Standard training plus access to Mission-Vet manual
 - GTO = planning, implementation (supervision, monitoring...), self evaluation (audit and feedback)
- **3 large VAMCs**
 - Case managers (69) randomized to IAU or IAU+GTO
 - 1500-2000 Veterans
- **RE-AIM measures**
 - Adoption = meeting 50% of eligible Veterans involved in intervention
 - Effectiveness = SUD, MH symptoms, functioning, housing

More Type 3 examples

Bauer et al. *Implementation Science* (2016) 11:22
DOI 10.1186/s13012-016-0385-7

Implementation Science

Kilbourne et al. *Implementation Science* 2014, 9:132
<http://www.implementationscience.com/content/9/1/132>



STUDY PROTOCOL

Open Access

Partnering with health system operations leadership to develop a controlled implementation trial



Mark S. Bauer^{1*}, Christopher Miller¹, Bo Kim¹, Robert Lew², Kendra Weaver³, Craig Coldwell⁴, Kathy Henderson⁵, Sally Holmes⁶, Marjorie Nealon Seibert⁶, Kelly Stolzmann⁶, A. Rani Elwy⁶ and JoAnn Kirchner⁵

Swindle et al. *Implementation Science* (2017) 12:90
DOI 10.1186/s13012-017-0624-6

Implementation Science

STUDY PROTOCOL

Open Access

Protocol: Adaptive Implementation of Effective Programs Trial (ADEPT): cluster randomized SMART trial comparing a standard versus enhanced implementation strategy to improve outcomes of a mood disorders program

Amy M Kilbourne^{1,2*}, Daniel Almirall³, Daniel Eisenberg⁴, Jeanette Waxmonsky^{5,6}, David E Goodrich^{1,2}, John C Fortney⁷, JoAnn E Kirchner^{8,9}, Leif I Solberg¹⁰, Deborah Main¹¹, Mark S Bauer¹², Julia Kyle^{1,2}, Susan A Murphy^{3,3}, Kristina M Nord^{1,2} and Marshall R Thomas^{5,6}

Cucciare et al. *Implementation Science* (2016) 11:65
DOI 10.1186/s13012-016-0432-4

Implementation Science

STUDY PROTOCOL

Open Access

A mixed methods protocol for developing and testing implementation strategies for evidence-based obesity prevention in childcare: a cluster randomized hybrid type III trial



Taren Swindle^{1*}, Susan L. Johnson², Leanne Whiteside-Mansell³ and Geoffrey M. Curran³

Cucciare et al. *Implementation Science* (2016) 11:65
DOI 10.1186/s13012-016-0432-4

STUDY PROTOCOL

Open Access

Assessing fidelity of cognitive behavioral therapy in rural VA clinics: design of a randomized implementation effectiveness (hybrid type III) trial



Michael A. Cucciare^{1,2,3,4*}, Geoffrey M. Curran^{1,2,5}, Michelle G. Craske⁶, Traci Abraham¹, Michael B. McCarthur¹, Kathy Marchant-Miros¹, Jan A. Lindsay^{3,4,7,8}, Michael R. Kauth^{3,4,7,8}, Sara J. Landes^{1,2,3,4} and Greer Sullivan⁹

Newish thinking on hybrid designs

- Changing thinking on “lack of fixed-ness” of interventions contributing to changing views on when and why of hybrid-type designs
- Hybrid type 1 less of a “special case” but more routine?
 - If effectiveness research is the “last step” before trying to get people to do the thing... why not more of a focus on implementation questions?
- Some folks doing hybrid 1 type work in efficacy research
- Type 2 designs need to be fully justified and include “failsafe”
 - Make scientific premise argument based on evidence of intervention *and* strategies
 - Clarity around intervention/strategy components essential
- Hybrid type 3 less of a “special case” also?
 - When wouldn’t we want patient-level outcomes data?
 - Clearly some tho... like perhaps uptake of vaccines
 - Shouldn’t we PROVE how much fidelity is important and under what circumstances?

What problems do people run into in trying to get hybrid studies funded?

- Disagreements over “how much evidence is enough” to begin including implementation focus
 - “But, we have no trials among people with green eyes...”
 - “Enough already! Get people to do the darn thing.”
- What if interventionist and/or context is REALLY different than in the effectiveness trials? (so more *major* adaptation expected)
 - LMIC research
 - How different is *too* different for hybrid?
- Not enough data on barriers/facilitators to uptake to ground selection of proposed implementation strategies
- No pilot data on implementation strategy (type 3)

What about blending EBPs?

- This is common...
- It's all about *“making the case”*
 - What is the reasonable action given the state of the evidence?
 - Think about why patients *do/do not* need to be randomized
- Each EBP has it's own strong evidence?
 - Package is new “thing”
 - “no reason to expect it won't work, but...” (type 3)
- Some have strong evidence some don't?
 - “Given the mixed data, we will actively collected clinical outcomes...”
 - Do clinical outcomes need to be “co-primary”? (type 2)

Worksheet to help decide...; Q 1

Effectiveness-Implementation Hybrid Designs D&I Workshop 2019

Geoffrey M. Curran, Sara J. Landes

Workshopping your idea: Questions to consider

- What is the nature of the effectiveness data on “the thing”?
 - “Very/pretty darn strong”, especially if not a lot of intervention adaptation needs to take place? Consider type 3 or type 2 depending on how much you also know about implementation factors (see below).
 - Mixed results? Missing (strong) effectiveness data? Consider Types 1 and 2.

Q2 and Q3

- How much do you expect “the thing” will need to be adapted for where you want to study/use it?
 - A little? Consider including adaptation process as a step in an implementation-focused project (so, more type 2 or 3).
 - A lot? Consider focusing on effectiveness in a type 1 or type 2 “pilot of implementation strategy” version.
- How much do you already know about the barriers/facilitators to the implementation of the thing in your context of interest?
 - Not much? If you also need to focus on effectiveness data right now, consider type 1.
 - Do you know enough already to develop and pilot test a “new” implementation strategy (or package)? Consider type 2 to “pilot” strategy.

Q4 and Q5

- Have you or someone else already tried an implementation strategy (or package) with your intervention and you know it didn't work very well?
 - If your intervention already has strong effectiveness data, this could be a great place to start for a type 3 or type 2 “pilot” for a new strategy (or package).
- Do you already have pilot data on a strategy (or package) that you want to test in a comparative study?
 - If your intervention effectiveness data are strong, consider a type 3.
 - If your intervention effectiveness data are mixed, consider a dual-randomized type 2.

Q6

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Q6

- After answering all of the above questions, do you still want to consider a hybrid design at all?
 - No? Feel free to run screaming from the room. We understand.
 - Yes? Seek out and learn from published protocol papers and other manuscripts describing studies that seem to be like what you want to do. Talk with people already funded to do the type of study you want to do. Talk with project officers/portfolio managers. Give us a call.

Thanks to these great folks for their thoughts and contributions to the work:

- Brian Mittman, PhD
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- JoAnn Kirchner, MD
- John Fortney, PhD

Question, comments, heckling...

