Grant Writing 101

“EXAMPLE: The American Heart Association"

Brian R. Wamhoff, Ph.D.

wamhoff@virignia.edu

Dept of Medicine, Cardiovascular Division
Other mechanisms:

• NIH – K99 Award
• JDRF
• ADA
• APS
• NRSA
• ACS
“Research broadly related to cardiovascular function and disease, stroke or related to clinical, **basic science**, bioengineering or biotechnology, and public health problems.”
"Optimizing your funding success with the American Heart Association"

1. My personal experiences with the AHA

2. The AHA Mid-Atlantic and National Affiliates

3. How the Review Process Works
   a. Reviewing the Pre- and Post-Doctoral Fellowship and *Grantsmanship Tips*
   b. Reviewing the Scientist Development Grant & the Beginning Grant-in-Aid

4. Concluding remarks
My personal experiences with the AHA

Pre-doctoral Fellowship

Post-doctoral Fellowship (declined for APS fellowship)
2002 – 2004 AHA Mid-Atlantic Affiliate, “Molecular mechanisms of decreased smooth muscle differentiation marker expression associated with the pathophysiology of atherosclerosis.” University of Virginia, PI: Gary K Owens, Ph.D.

Scientist Development Grant (accepted), Beginning Grant-in-Aid (rejected)

Used SDG to generate data for current NIH RO1 – 07/01/06 – 06/31/11

Study Section
2005 –  AHA National Affiliate BASIC 1 Study Section

Board of Directors
2005 –  AHA Local Affiliate, spokesperson for AHA impact on UVA funded research and training young cardiovascular scientists
The AHA Mid-Atlantic and National Affiliates

http://www.americanheart.org/presenter.jhtml?identifier=10813

**Mid-Atlantic Affiliate**

**Deadline:**
Jan, 2007

**Offering Programs:**
*Pre-doctoral* (29/115 25% down from 34%)
*Post-doctoral* (21/99 21% up from 15%)
*Beginning Grant-in-Aid* (22/74 30% up from 14%)

**National Affiliate** also collectively reviews National, Greater Midwest, Heartland and Pacific Mountain Affiliates

**Deadline:**
July, 2006
Jan, 2007

**Offering Programs:**

*Scientist Development Grant* (85/356 24%, $65K, 3-4y)

Fellow-to-Faculty
NIH paylines are dropping at an astounding rate!

Current pay lines are <12% for NHLBI, down from approximately 20-24% in 2004-05

**SOLUTION:** Start writing good grants early in your career because this is what you will be doing for the rest of your career, in good times and bad!

Post-docs that have funding keep their jobs.
Current AHA Funded Proposals at CVRC

**Mid-Atlantic Affiliate**

Pre-doctoral fellows ($20K 2+1y):
Jeremy Mauldin
James Thomas
Matt Alexander

Post-doctoral fellows ($35K 2+1y):
Elena Galinka
Tracy Deem
Anthony Ore

Beginning Grant-in-aid ($66K 2y):
Elaine Felecia Etter
Brant Isakson

**National Affiliate**

Scientist Development Grant ($66K, 4y):
Brian Wamhoff
Tadashi Yoshida
How the Review Process Works
“Prior to Study Section”

Proposals are assigned to a study section based on the specific “codes” you select for describing your grant.
Science Focus

“Research broadly related to cardiovascular function and disease, stroke or related to clinical, basic science, bioengineering or biotechnology, and public health problems.”
The proposal does not need to be related to CVD!

Mid-Atlantic Affiliate

**Mid-Atlantic 1A**
Integrative Cardiac Biology/Regulation  
Radiology & imaging  
Surgery

**Mid-Atlantic B**
Electrophysiology & Arrhythmias/Regulation  
Vascular Biology & Blood Pressure/Regulation  
Cardiovascular Regulation (autonomic regulation)

**Mid-Atlantic 2**
Lipoproteins & Lipid Metabolism  
Thrombosis  
Vascular Wall Biology

**Mid-Atlantic 3**
Cardiorenal  
Lung, Respiration & Resuscitation  
Immunology & Microbiology

**Mid-Atlantic 4**
Cell Transport & Metabolism  
Cellular CV Physiology & Pharmacology

**Mid-Atlantic 5**
Molecular Signaling

**Mid-Atlantic 6**
Basic Cell & Molecular Biology  
CV Development
The proposal does not need to be related to CVD!

National Affiliate

Basic Cell & Molecular Biology 1 (25 members)
  Basic Cell & Molecular Biology 2
  Behavioral Science, Epidemiology & Prevention
  Bioengineering & Biotechnology
  Brain
  Cardiorenal
  Cardiovascular Development
  Cardiovascular Medical Research and Education Fund
  Cell Transport Function & Metabolism/Electrophysiology & Arrhythmias
  Immunology & Microbiology
  Integrative Cardiac Biology/Regulation
  Lipoproteins, Lipid Metabolism & Nutrition
  Lung, Resuscitation & Respiration
  Molecular Signaling 1
  Molecular Signaling 2
  Radiology, Imaging & Surgery
  Thrombosis
  Vascular Biology & Blood Pressure/Regulation
  Vascular Wall Biology 1
  Vascular Wall Biology 2
How the Review Process Works
“Prior to Study Section”

*Putting yourself in the mindset of the Reviewer can only help.*

Applications are received 1 month prior to study section.

The average time spent reviewing an application is **2.5-4 hrs**.

Each Reviewer receives 12-14 applications: **12 x 3 = ~36 hrs**.

For each application, the Reviewer is assigned as:
- Primary Reviewer (R1)
- Secondary Reviewer (R2)
- Reader (R3)

The Reviewer critiques and scores all applications **collectively** according to AHA guidelines (Scale of 1-5, where 1 is outstanding)

33% of all applications are **streamlined**.

All scores are posted online 1 week prior to study section.
**OBJECTIVE:** Score applications using the full range of scores

**SAMPLE:**
Reviewer scored 10 applications = (7 primary or secondary assignments with 3 reader assignments)

**PRIORITY SCORE RANGE:** (Below is an example of an appropriate distribution of scores)

<table>
<thead>
<tr>
<th>1.0</th>
<th>1.5</th>
<th>2.0</th>
<th>2.5</th>
<th>3.0</th>
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<td>2</td>
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<td>1</td>
<td>1</td>
<td>0</td>
<td>1</td>
<td></td>
</tr>
</tbody>
</table>

**Note:** all pre-doc, post-doc, SDG and BGIA are scored collectively

**SAMPLE DISTRIBUTION OF APPLICATIONS AMONG PRIORITY SCORE INTERVALS:**
(Reviewer should evaluate distribution for use of full range prior to posting critiques and scores on the Electronic Peer Review website)

<table>
<thead>
<tr>
<th>1.0</th>
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<th>2.5</th>
<th>3.0</th>
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<td></td>
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</tbody>
</table>

**STREAMLINED APPS = 33%**
Pre-doctoral & Post-doctoral Fellowships

**Objective**: To help students or trainees initiate careers in cardiovascular and stroke research by providing research assistance and training. **Awardees will be expected to devote full time to research or activities directly related to their development into independent researchers.**

**CRITIQUE LAYOUT**

A. **Name of Applicant & Research Project Title**

B. **Proposal Description**: Briefly describe the project, (approximately one paragraph) including:
   - Scientific objectives
   - Research design and methods

C. **Critique**: Address the following criteria in this order, listing strengths and weaknesses include detailed suggestions for enhancements for a future resubmission. **Use the quality of these areas as comparison points for assigning priority scores to applicants in these programs:**

1. **Evaluation of Proposal**
   - (1/3rd of score): 1.3
   - Location within application:
     - Body of 12 page scientific proposal

   - Scientific merit of proposed project
   - Probability of achieving scientific objectives

2. **Evaluation of Investigator**
   - (1/3rd of score): 1.9
   - Location within application:
     - Page 4 and quality of scientific proposal
     - Pages 4-7-8, biosketch and page 11
     - Pages 4-7-8, biosketch, other support page and any included reprints from applicant
     - Listed references/ bibliography
     - Pages 4-7-8 and biosketch
     - Letters included in application documents, names on page 10

3. **Evaluation of the Sponsor/Institution**
   - (1/3rd of score): 2.5
   - Location within application:
     - Sponsors information pages, other support page
     - Sponsors information pages
     - Sponsors information pages
     - Body of 12 page scientific proposal, Sponsors information pages – detailed training program
     - Sponsors information pages, other support page

   **A. Qualifications of the sponsor: Sponsor’s reputation as an investigator**
   - Publications and funding history
   - Track record with trainees
   - Commitment to applicant’s training
   - Quality and comprehensiveness of the Training plan
   - Current funding available to support Fellow’s project

   **B. Qualification of the Facilities and resources available from institution**
   - Space stated in the sponsor’s laboratory
   - Evidence of exposure to a research training environment.

**1.9 (1.8-2.0)**
1. The **SCIENCE** must be **good**!

2. Simple, clear transition of thought process, structure. **Simple, Simple, Simple.**

3. Have a **HYPOTHESIS**.

4. **FOCUSSED**, not **OVERAMBITUOUS**

   #1 comment of reviewers: “**This is grant is unfocused and overambitious.**”

5. Mechanistic not Descriptive.

   #2 comment of reviewers: “**This Aim is descriptive.**”

6. Clearly state **CAVEATS** and **POTENTIAL PITFALLS** for each Aim.

7. Supportive **preliminary data**, whether it’s your data, data from a previous member of your lab or data from another lab.
APPLICATION FOR RESEARCH AWARD

PROJECT SUMMARY

1. Applicant:
   Brian R. Warnhoff
   Ph.D.

2. Institution where work will be done:
   University of Virginia

3. Sponsor (if applicable):
   Gary K. Owens
   Ph.D.

4. Department Head:
   Andrew P. Sonty
   M.D.

5. Applying to:
   Mid-Atlantic Affiliate

6. Award program:
   Postdoctoral Fellowship
   Resubmission?
   Yes

7. Dates of proposed award:
   Start: 07/01/2002
   End: 06/30/2004

8. Project title (limit to 120 characters or less):
   Molecular mechanisms of decreased smooth muscle differentiation marker expression associated with atherosclerosis.

9. Project summary (must be completed on this page):
   Atherosclerosis, the principal cause of heart attack and stroke, is a complex inflammatory-fibroproliferative response to various forms of insult to the endothelium resulting in phenotypic modulation of smooth muscle cells (SMC) of the artery wall (FIG 1). A characteristic feature of SMC phenotypic modulation is decreased expression of SMC differentiation marker proteins (i.e., smooth muscle myosin heavy chain (SM-MHC) and SM22alpha), increased proliferation, decreased contractility and increased matrix production - that contribute to lesion formation. The overall aim of this proposal is to determine molecular mechanisms that contribute to the decreased SMC-specific differentiation marker expression associated with vascular injury and atherosclerosis. Recent studies from our lab were the first to show that transcriptional regulation of the SM-MHC and SM22alpha promoter are decreased in transgenic mice following carotid injury (FIG 3). We identified a "novel mechanism" of SM22alpha gene regulation whereby mutation of the SM22alpha G/C-rich repressor cis regulatory element attenuated the injury-induced decrease of the SM22alpha transcript. Further, we showed by electrophoretic mobility shift assays that the transcription factors Sp1 and Sp3 (both of which are increased in SMCs in models of vascular injury) bind to the G/C-rich repressor region (FIG 4). However, it is UNKNOWN whether 1) Sp1/Sp3 or other transcription factors contribute to the repressor effect in vivo and 2) how SM-MHC and SM22alpha gene expression are regulated in an animal model of atherosclerosis (TABLE 1). Therefore, my overall hypothesis is that decreased SM-MHC and SM22alpha gene expression associated with atherosclerosis is mediated, in part, by Sp1/Sp3 binding to the G/C repressor region of the SM-MHC and SM22alpha genes. This hypothesis will be tested by addressing the following three specific aims: 1) To determine the role of the G/C repressor in regulating SM-MHC and SM22alpha gene expression in the ApoE-/- mouse model of experimental atherosclerosis with and without vascular injury; 2) To determine trans-acting factors and mechanisms that regulate the activity of the SM-MHC and SM22alpha G/C repressor elements. Including the role Sp1 and Sp3; 3) To determine the ability of the factors identified in Aim 2 to modulate injury-induced SMC gene expression in vivo using adenoviral-mediated gene transfer. The experimental design will integrate cellular and molecular biology with animal models to address the overall hypothesis.

10. Amount Requested:
   Year 1: $30,000
   Year 2: $30,000
   Year 3: $0
   Year 4: $0
   Year 5: $0

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Page 1
Layout of the proposal

**Project Summary** (250-300 word Abstract)

Phenomena X or disease X is… A characteristic feature of this process is… Although ABC has been shown to… it is unknown whether… **Preliminary studies** [or **Recent studies** from our lab] show that… However, it is unknown whether… Therefore, the **overall hypothesis** is that… This hypothesis will be tested by the following **specific Aims:** Aim 1 will determine… Aim 2 will determine… Aim 3 will determine…

A. **Specific Aims** (1 page)

B. **Background** (3 pages)

C. **Research Design and Methods** (7.5 pages)

D. **Ethical Aspects of Proposed Research** (1/2 page)
A. **Specific Aims** (1 page)

This Page will “make” or “break” your application. When the reviewer has finished reading this page, they will already have a preconceived notion of the quality of the proposal and a score.

[Restate your original Project Summary and expand on each Aim]

Phenomena X or disease X is… A characteristic feature of this process is… Although ABC has been shown to… it is unknown whether… **Preliminary studies** [or **Recent studies** from our lab] show that… However, it is unknown whether… Therefore, the **overall hypothesis** is that… This hypothesis will be tested by the following **specific Aims**:

**Aim 1** will determine… **Aim 1** will utilize X and Y methodology to… In **Aim 1A** we will… In **Aim 2A**… We hypothesize that…

**Aim 2** will determine…

**Aim 3** will determine…

The **results of this study** will lead to a better understanding of….
Tips on Aims

1. Your aims should be **interconnected but not dependent** on the successful outcome of another aim.

   **EXAMPLE:**  
   **Bad** – Aim 2 cannot proceed until the studies in Aim 1 are completed.  
   **Good** – Aim 2 proceeds in parallel with Aim 1 and findings from Aim 1 might direct future studies in Aim 2 or 3.

   In the end, *aims relate back to the overall hypothesis.*
Tips on Aims

2. If the Aims are not interconnected, the project can be perceived as “overambitious and unfocussed” where each Aim is probably a proposal in itself.

3. If you cannot keep your Aims page to 1-1.5 pages, then you are proposing too much and the grant is probably “overambitious and unfocussed”.

4. If the project is 2 years, then the probability of achieving the Aims should be 2 years. Proof that the applicant has thought this through is usually addressed in Section C, Predicted Results/Interpretation of Results and with a timeline or timeline statement.

5. The standard rule of thumb for a pre/post-doc fellowship is two Aims. It is OK to propose three Aims. However, if Aim 3 will not fit into the 2 year timeline, but it is clearly a logical progression of the studies, then simply state:

   *Aim 3 is to determine the... Although this Aim does not fit the time frame of this proposal, future studies by the applicant will...*

6. **Descriptive Aims:** If the Aim cannot have a stand alone hypothesis, then it is probably “descriptive”, not “mechanistic”, and may be detrimental to the success of the grant. **Example:** gene arrays (this ties into pt 1).
Tips on Aims

7. **Never propose to make a knockout mouse or transgenic mouse** for a 2 year proposal. If you do not have the mouse in-house, you are not ready to submit a proposal. These proposals are viewed as risky, especially during tight funding periods. If you have the mouse in-house, show preliminary data.

   For example, you received or made a mouse null for XYZ. Show a Southern blot with the XYZ deletion or histology images that show a phenotype, etc…

8. **Developing a new technology is risky.** For example, if you are proposing to measure flow patterns in diseased blood vessels but Aim 1 is to complete the technology, this will probably not get funded.

   Use tools, models, animals that are readily available to you.
B. Background (3 pages)

1. **Do not assume that the reviewer is an expert in your field!**

2. Expand on the brief background that has already been stated in the Project Summary. *Use the Project Summary as your outline* (subsections for Section B).

3. If the mechanisms you are proposing are complex on paper and thus very difficult to visualize in one’s mind, make a **Schematic/Cartoon** that you can refer to throughout the proposal, in your aims and in your predicted results.

   For example: ABC regulates XYZ. Although we propose ABC regulates XYZ via 1, 2, and 3 (Aim 1), 1 can also activate 4 and 5 to regulate XYZ (Aim 1a). Moreover, preliminary studies show that ABC can mediate XYZ via 6 (Aim 2).
Figure 1. Overall hypothesis for this proposal.
4. Avoid jargon and multiple abbreviations, e.g. **VGCC mediates Ca influx to activate ROK-dependent activation of SMGX.** VGCC = voltage-gated Ca channel, Ca = calcium, ROK = Rho kinase, SMGX = smooth muscle cell gene expression.

Use abbreviations for terms that are used throughout the proposal and are obvious.

5. Preliminary data: Although *preliminary data are not required for a pre-doc*, **show preliminary data.** Preliminary data may simply be proof that you can do the exps your proposing or that a critical exp has been performed by another lab or someone previously in your lab. However, if there is a key piece of data that your overall hypothesis hinges on, you must show that data.

For example, if you are proposing that ABC effects XYZ by 123. You should have the preliminary data showing that ABC effects XYZ. Each Aim will then determine 123.

![Diagram showing ABC effects XYZ](image)

You may only have 1 preliminary Figure that is yours.

***6. Show the reviewer the experiment is feasible even if the data do not address the specific hypothesis – PROOF OF PRINCIPLE DATA!***
C. Research Design and Methods (7.5 pages)

Restate the overall hypothesis. Keep the Reviewer focused.

**Specific Aim 1:** To determine...

**Rationale:**
Briefly restate why you're doing this aim. The hypothesis for Aim 1 is that...

**Experimental Design:**
Unless absolutely necessary to the question being asked, you do not need details of the experiment that include pH of solutions, time of transfection, how RNA is isolated, etc.

**Interpretation of Results:**
1) State what you predict will happen.  2) State what can go wrong and how you will interpret these findings. This is critical and shows the reviewer that you have thought through all of the experimental parameters and outcomes. Have alternative hypotheses. As your mentor has probably said, “99% science is failure and the 1% success is learning from failure” – every reviewer knows this.

**Future Directions:**
If there are future directions beyond the scope of this proposal/timeframe, briefly state that you are aware of this; a sign that you see beyond the limited scope of this proposal.

**Timeline:** End Section C with a timeline or course of action for each Aim over 2 years.
D. Ethical Aspects of Proposed Research (1/2 page)

For example, if you are using animals, is it absolutely necessary or can these same questions be addressed *in vitro*?

Modified from Wamhoff, post-doc: Although *in vitro* studies can provide substantial information regarding the molecular mechanisms regulating of X, such data may be confounded by the changes that occur when the cell is not in its native *in vivo* setting. Thus, in addition to detailed *in vitro* studies, it is essential to assess the results of key genetic manipulations in an integrated manner in terms of organ/whole animal phenotype and there is no alternative to studies in animals. The mouse is the mammalian model of choice for studies of genetic manipulation due to its small size, rapid breeding, low costs and the swiftly increasing knowledge of its genome.

**Cell culture vs. animals**
<table>
<thead>
<tr>
<th>2. Evaluation of Investigator: (1/3rd of score):</th>
<th>Location within application:</th>
</tr>
</thead>
<tbody>
<tr>
<td>Evidence of potential for a research career</td>
<td>Page 4 and quality of scientific proposal</td>
</tr>
<tr>
<td>Trainee’s academic record (academic achievement, GPA)</td>
<td>Pages 4-7-8, biosketch and page 11</td>
</tr>
<tr>
<td>Quality of trainee’s previous research</td>
<td>Pages 4-7-8, biosketch, other support page and any included reprints from applicant</td>
</tr>
<tr>
<td>Trainee’s familiarity with literature and work of other investigators in the field of interest</td>
<td>Listed references/bibliography</td>
</tr>
<tr>
<td>Trainee’s need for the training experience</td>
<td>Pages 4-7-8 and biosketch</td>
</tr>
<tr>
<td>** Quality of required 3 reference letters (do not use specific names)</td>
<td>Letters included in application documents, names on page 10</td>
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</table>

This is YOU

Reference letters are CRITICAL!

Get people who **know** you as a **scientist** and **person** to write your letters.

Example, if you tanked a few classes as an undergraduate, have someone write a letter that emphasizes how you’ve changed since then, if applicable 😊
This is your PI and UVA

The PI’s *training plan* is CRITICAL!

Can and how will the PI and the PI’s environment turn you into a world-class scientist?

1. Sponsor’s research and applicant’s connection to this work.
2. Sponsor’s plan to develop the applicants research capabilities and a sequence in which the applicant will be given responsibility to conduct the research.
3. Indicate other training or course work required for this proposal.
4. Relationship of the research training plan to your career goals, i.e. does your PI have any clue what you want to be in the future?

Your PI must be able to currently fund your work.
Read your PI’s training plan before submitting. Make sure it addresses all 4 points.
**Pre-doctoral & Post-doctoral Fellowships**

**Objective:** To help students or trainees initiate careers in cardiovascular and stroke research by providing research assistance and training. *Awardees will be expected to devote full time to research or activities directly related to their development into independent researchers.*

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**CRITIQUE LAYOUT**

A. Name of Applicant & Research Project Title

B. Proposal Description: Briefly describe the project, (approximately one paragraph) including:
   - Scientific objectives
   - Research design and methods

C. Critique: Address the following criteria in this order, listing strengths and weaknesses include detailed suggestions for enhancements for a future resubmission. *Use the quality of these areas as comparison points for assigning priority scores to applicants in these programs*

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<td>Scientific merit of proposed project</td>
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<tr>
<td>Probability of achieving scientific objectives</td>
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<tbody>
<tr>
<td>Evidence of potential for a research career</td>
<td>1.9</td>
<td>Page 4 and quality of scientific proposal</td>
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<thead>
<tr>
<th>3. Evaluation of the Sponsor/Institution:</th>
<th>(1/3)rd of score:</th>
<th>Location within application:</th>
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</thead>
<tbody>
<tr>
<td>Qualifications of the sponsor: Sponsor’s reputation as an investigator</td>
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<td>Sponsors information pages, other support page</td>
</tr>
<tr>
<td>Publications and funding history</td>
<td></td>
<td>Sponsors information pages</td>
</tr>
<tr>
<td>Track record with trainees</td>
<td></td>
<td>Sponsors information pages</td>
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<tr>
<td>Commitment to applicant’s training</td>
<td></td>
<td>Sponsors information pages</td>
</tr>
<tr>
<td>Quality and comprehensiveness of the Training plan</td>
<td>1.6</td>
<td>Body of 12 page scientific proposal, Sponsors information pages – detailed training program</td>
</tr>
<tr>
<td>Current funding available to support Fellow’s project</td>
<td></td>
<td>Sponsors information pages, other support page</td>
</tr>
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</table>

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<thead>
<tr>
<th>B. Qualification of the Facilities and resources available from institution:</th>
<th>Location within application:</th>
</tr>
</thead>
<tbody>
<tr>
<td>Space stated in the sponsor's laboratory</td>
<td>Page 11 &amp; 12</td>
</tr>
<tr>
<td>Evidence of exposure to a research training environment.</td>
<td>Page 12, reviewer knowledge of institution</td>
</tr>
</tbody>
</table>

1.6 2.1 3.0
**Scientist Development Grant**

**Objective** - To support highly promising beginning scientists in their progress toward independence by encouraging and adequately funding research projects that can bridge the gap between completion of research training and readiness for successful competition as an independent investigator. At the time of award activation, no more than four years will have elapsed since an applicant's first faculty/staff appointment at the assistant professor level or its equivalent. A pivotal requirement is the demonstration that the award will promote independent status for the applicant. An applicant cannot hold, nor have held, any other national award prior to the activation date of the Scientist Development Grant. A sponsor should not be required, although it is important that applicant’s Department Head provides assurance that the applicant has the institution’s support.

**CRITIQUE LAYOUT**

**A. Name of Applicant & Research Project Title**

**B. Proposal Description:** Briefly describe the project, (approximately one paragraph) including:
- Scientific objectives;
- Research design and methods.

**C. Critique:** Address the following criteria in this order, listing strengths and weakness include detailed suggestions for enhancements for a future resubmission. Use the quality of these areas as comparison points for assigning priority scores to applicants in these programs.

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<th>1. Future Independence of Investigator:</th>
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<tr>
<td>Demonstrated evidence that the award will promote independent status for the applicant by the end of the 3 or 4-year award. Should not serve as enhanced funding for professional personnel working on the research program of an established scientist.</td>
<td>Mentioned in SDG reference letters and department head letters</td>
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<tbody>
<tr>
<td>Originality of ideas</td>
<td>Body of 12 page scientific proposal, mention in reference letters</td>
</tr>
<tr>
<td>Projected scientific impact</td>
<td>Body of 12 page scientific proposal, mention in reference letters</td>
</tr>
<tr>
<td>Soundness of approach</td>
<td>Body of 12 page scientific proposal</td>
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<td>Logical organization</td>
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<th>3. Investigator - Qualifications of the applicant:</th>
<th>Location within application:</th>
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<tr>
<td>Appropriate professional career stage</td>
<td>Pages 3, 4, and 7, Biosketch</td>
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<td>Pages 4, 7, 14 and Biosketch</td>
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<td>Current relationship to supervisor/mentor</td>
<td>Might be mentioned in Dept. Head Letter, page 15, or in reference letters</td>
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<td>Prior awards, if any, and resulting accomplishments</td>
<td>Pages 4, 7, 14 and Biosketch</td>
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<td>Number and quality of publications in peer reviewed journals</td>
<td>Biosketch</td>
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<td>Quality of required 3 reference letters (do not use specific names)</td>
<td>Page 10 lists names, 3 referent letters</td>
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<th>4. Qualification of the Facilities and resources available from institution:</th>
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<td>Adequacy of available resources, including consultative support, and facilities</td>
<td>Pages 11-14</td>
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<tr>
<td>Institutional commitment as demonstrated in Department Head letter</td>
<td>Pages 6, 11 and 15</td>
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Beginning Grant-in-Aid

Objective - To promote the independent status of promising beginning scientists. At application, applicant must be an M.D., Ph.D., D.O., D.V.M. or equivalent initiating independent research career. Applicant must have faculty/staff appointment (up to and including assistant professor or equivalent) at activation. A sponsor should not be required, although it is important that applicant’s Department Head provides assurance that the applicant has the institution’s support.

CRITIQUE LAYOUT

A. Name of Applicant & Research Project Title

B. Proposal Description: Briefly describe the project, (approximately one paragraph) including;
   - Scientific objectives
   - Research design and methods

C. Critique: Address the following criteria in this order, listing strengths and weaknesses include detailed suggestions for enhancements for a future resubmission. Use the quality of these areas as comparison points for assigning priority scores to applicants in these programs.

1. Future Independence of Investigator:

   | Demonstrated evidence that the award will promote independent status for the applicant at the completion of first or second award. Should not serve as enhanced funding for professional personnel working on the research program of an established scientist. | Location within application: Mentioned in department head letter |

2. Scientific excellence of the research proposal:

   | Originality of ideas | Location within application: Body of 12 page scientific proposal |
   | Projected scientific impact | Body of 12 page scientific proposal |
   | Soundness of approach | Body of 12 page scientific proposal |
   | Logical organization | Body of 12 page scientific proposal |

3. Investigator - Qualifications of the applicant:

   | Appropriate professional career stage | Location within application: Pages 3, 4, and 7, Bioresearch |
   | Relevant experience | Pages 4, 7 and Bioresearch |
   | Productivity in previous training and research | Pages 4, 7, 14 and Bioresearch |
   | Current relationship to supervisor/mentor | Might be mentioned in Dept Head Letter |
   | Prior awards, if any, and resulting accomplishments | Pages 4, 7, and Bioresearch |
   | Number and quality of publications in peer reviewed journals | Bioresearch |

4. Qualification of the Facilities and resources available from institution:

   | Adequacy of available resources, including consultative support, and facilities | Location within application: Pages 11 - 14 |
   | Institutional commitment as demonstrated in Department Head letter | Pages 6, 11 and 15 |
The SDG and BGA are very similar. What these grants ultimately come down to are “independence issues”:

1. The science must be excellent, good grantmanship, feasibility, etc.
2. Will the award foster independence and National funding?

Classic situation: The applicant still resides at the institute where they trained as a postdoctoral fellow. In fact, the applicant most likely has dedicated lab space within the former PI’s lab (at least on paper). Is the work proposed in this application different enough from the former PI that this work will eventually lead to funding on the national level, i.e. NIH RO1?

The #1 Reviewer Comment: I am not convinced that the applicant is/will be independent from their previous mentor.

What convinces a reviewer of independence or strive to be independent:

1. Department Chair letter and commitment to space (non-issue for tenure track).
2. Letter from former mentor clearly stating how this work is different.
3. Prior awards and quality of publications and how they relate to this proposal (not necessarily quantity).
How the Review Process Works  
“Day of Study Section”

**LENGTH:** 1 day  
**START:** 7:00 AM  
**END:** 6:00 PM or when last app is discussed

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**STREAMLINED APPS = 33%**

Only if a proposal received 2 or more “streamlines” will it not be discussed.
1. Anyone with COI leaves.
2. R1, R2 and R3 state preliminary scores ranges, e.g. R1 “1.9-2.1”, R2 “1.6-1.9”, R3 “1.7-2.0”
3. R1 gives <5 min summary of grant and 2-3 min discussion of concerns (no more than 10 minutes!)
4. R2 only adds comments that differ from R1 or “in agreement”
5. R3 adds comments that differ from R1, R2 or “in agreement”
6. R1, R2, R3 restate range of scores.
7. Each reviewer writes down their score.
8. Next application.

TOTAL TIME FOR YOUR APP = 7-12 min!
You receive your score and critiques via email, What do you do now?

1. **Funded**: Jump for joy, push on.

2. **Not funded**: Throw a temper tantrum, blame the world, push on, resubmit!
A large percentage of first-time submissions to the AHA get rejected.

This will happen to you. Breath.

Read your critiques, address every comment and resubmit.

Briefly point out what the reviewers like about the grant and then address every comment.

Walk the reviewer through all of your changes by clearly denoting in the text where the changes were made. *It is likely you will only get 1 of the 3 previous reviewers.*

This is not the time nor place to pick a fight. However, if the reviewer is completely off-base, be respectful. “I respectfully disagree with this comment. It has been shown that…”

Do what the reviewers ask.
INTRODUCTION: This is the first resubmission of RO1 HL081682 by a First-time Investigator. The “Summary of Critique” for the original submission (Oct 1, 2004) was received on May 20, 2005. I received a score of ## and the funding payline was ##. The applicant has made significant progress to address the Reviewers’ comments and suggestions, providing substantial new preliminary data and greatly improving the focus and quality of this proposal. In general, the major revisions encompassed: 1) general experimental clarification and validation of preliminary studies, 2) “descriptive nature of Aim 3”, and 3) “independence of the investigator”.

We thank the Reviewers for their overall enthusiasm for this proposal: 1) “a highly original and innovative proposal”, 2) “employs state of the art methods… and superb models… to address important and novel questions… for understanding phenotypically modulated SMCs”, 3) “well crafted research design” and “well written”, 4) “theory that has clearly been under-investigated”, 5) “excellent group of collaborators” with an “appropriate” environment, and 6) “preliminary data that are supportive of each aim”.

However, Reviewer 1 and 2 had a major concern regarding the “descriptive nature of the third specific Aim”. We now provide new preliminary data in this resubmission that will allow us to apply “transgenic mouse methodology (for) in vivo studies” to “advance the work more appropriately and link the studies to the first two Aims” (below, pt. 3). Our overall enthusiasm was also tempered by several excellent concerns and needed clarifications regarding preliminary studies/models raised in Critique 2 and addressed below:

1. “It is not clear whether the applicant can target experiments towards relevant directions and can recognize findings that influence clinical therapy.” Although it is out of the scope of this proposal to immediately translate findings into a clinical therapy, the new advances we have made in such a short period of time show that we can recognize findings that test mechanisms towards relevant directions. On page 25…

2. “the conclusion that (S1P effects) are mediated through calcineurin are premature”. We have expanded our preliminary results implicating calcineurin as a downstream regulator of S1P mediated SMGX, not depolarization.
Mid-Atlantic and National Affiliate can be on opposite sides of the spectrum

**TITLE**: Calcium-dependent regulation of smooth muscle phenotype
**PI**: Brian R Wamhoff

**Mid-Atl BGIA**
Score: 2.575
Percentile Rank: 58.33
Major Comments:
*R1*: The experimental plan is not well focussed… and as a result extremely ambitious and lack direction.
*R2*: …will not do much to bolster independence from current PI…
*R3*: Clearly at stage for an independent award

**National SDG**
Score: 1.5182
Percentile Rank: 9.88
Major Comments:
*R1*: Reasonable approach to test the hypothesis and logically organized.
*R2*: Supervisor supports the full independence of the researcher.
*R3*: …rising star…exceptional environment at UVa…
Concluding Remarks

In the end, the best grants/science get funded, for the most part 😊

Have your peers, former and current AHA fellows read your Specific Aims page. Do they understand what you are proposing to do?

Accept criticism openly but know what to filter and what not to filter.

Keep it simple, organized, and structured in such a way that if a Reviewer does not understand what you are proposing to do, it is because you failed, not them 😊

Even if your grant is not funded, you now have a detailed plan for the next 2+ years of your career. Not many people can say that 😊
Acknowledgements

Mike Sturek, PhD, University of Missouri
Gary Owens, PhD, University of Virginia

The American Heart Association
The American Physiological Society
Pfizer
FEST/UVA
NIH