- -How to success
- -How to do research
- -How to write

How to success

Three rules

ตรงต่อเวลา

อิทธิบาท ๔

รู้คุณคาของงาน

การทำงาน

พระบรมราโชวาท พระราชทานเนื่องใน โอกาสวันข้าราชการพลเรือน

"...เมื่อทำงาน ต้องมุ่งถึง 1 เมษายน 2532 จุดมุ่งหมาย ที่แท้จริงของงาน

งานจึงจะสำเร็จ

ได้รับประโยชน์ครบถ้วน ทั้งประโยชน์ของงาน และประโยชน์ของผู้ทำ..."

http://www.artskru.com/main/rachor/work.html

Impact Factor vs True Impact

Why Impact Factor?

Number is easier to understand than Science.

For example: signal transduction

What is true impact study?

- Interesting Cloning
- Knowledge Signal Transduction
- Application A Green Fluorescent

New

Broad vs Specific Condition

Living organism – Eukaryote – Human – Race – Regional

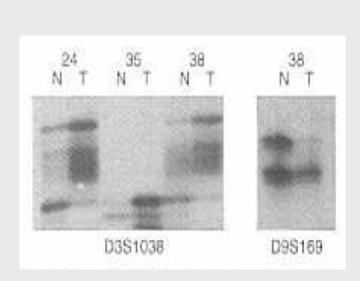
- p53 mutation in cancer
- p53 mutation in breast cancer
- p53 mutation in breast cancer in Thailand

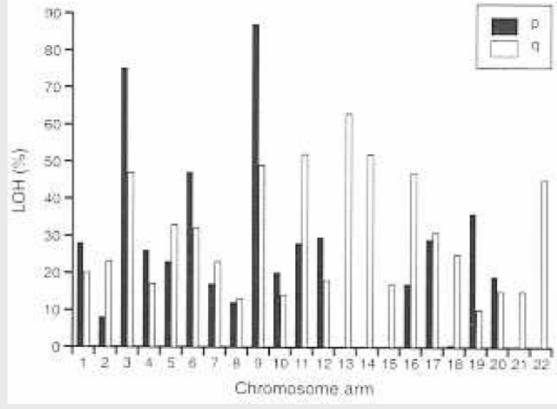
How to conduct research

•ตามหลังผู้ใหญ่ (อย่างสร้างสรรค์) หมาไม่กัด

The main article of my first TRF grant.

Mutirangura A, Tanunyutthawongese C, Pornthanakasem W, Kerekhanjanarong V, Sriuranpong V, Yenrudi S, et al. Genomic alterations in nasopharyngeal carcinoma: loss of heterozygosity and Epstein-Barr virus infection. Br J Cancer. 1997;76(6):770-6.

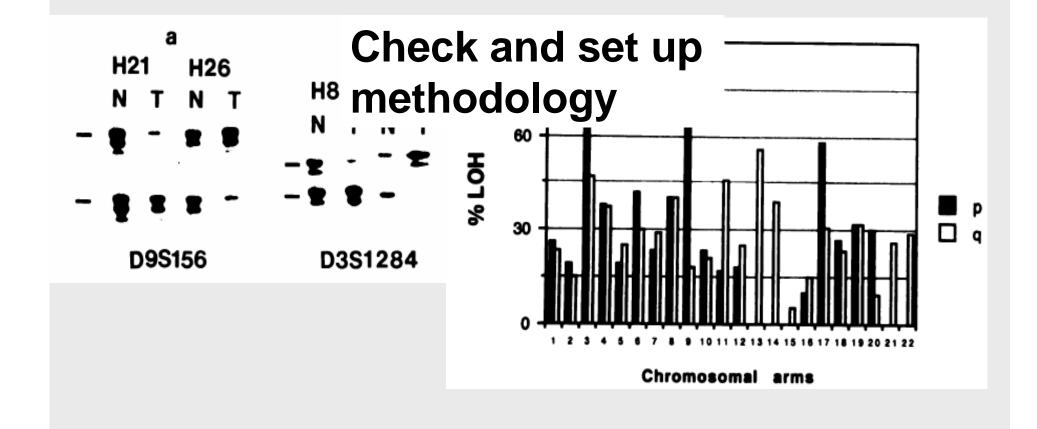




Allelotype of Head and Neck Squamous Cell Carcinoma¹

Homaira Nawroz, Peter van der Riet, Ralph H. Hruban, Wayne Koch, J. Michael Ruppert, and David Sidransky²

Department of Otolaryngology, Division of Head and Neck Cancer Research, Johns Hopkins University, Baltimore, Maryland 21205-2196 [H. N., P. V. D. R., W. K., J. M. R., D. S.], and Department of Pathology, Johns Hopkins Hospital, Baltimore, Maryland 21205 [R. H. H.]



How to conduct research

• จับแพะชนแกะ

การค้นพบโดยสมมุตติฐาน

ค้นพบ DNA ของไวรัสอีบีวี ในน้ำเหลืองของผู้ป่วยมะเร็งโพรงหลังจมูก

- 1. ไวรัส อีบีวี ใน มะเร็งโพรงหลังจมูก
- 2. DNA ของมะเร็ง ใน น้ำเหลือง (serum or plasma)



Tumor-specific DNA in

serum/plasma

Anker P et al Gastroenterology 1997, 112:1114-1120

Chen XQ et al Nat Med 1996, 2(9):1033-5

of cancer patients Nawroz H et al Nat Med 1996, 2(9):1035-7



Epstein-Barr viral DNA in serum/plasma of nasopharyngeal cancer patients

Mutirangura et al Clin Cancer Research 1998, 4:665-669



Quantitative EBV DNA in plasma of NPC patients as a tumor marker.

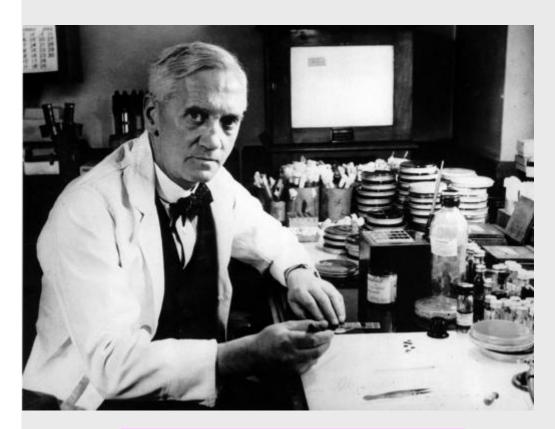
Lo YM et al. Cancer Res. 1999 Nov 1;59(21):5452-5.

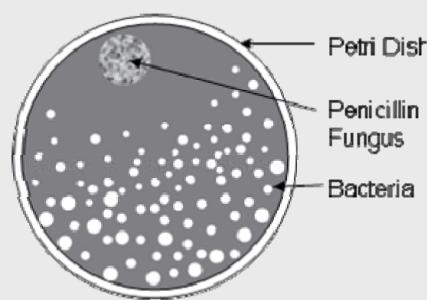
Cancer Res. 1999 Mar 15:59(6):1188-91.

How to conduct research

• โดยบังเอิญ

Serendipitous Discovery (การค้นพบโดยบังเอิญ)

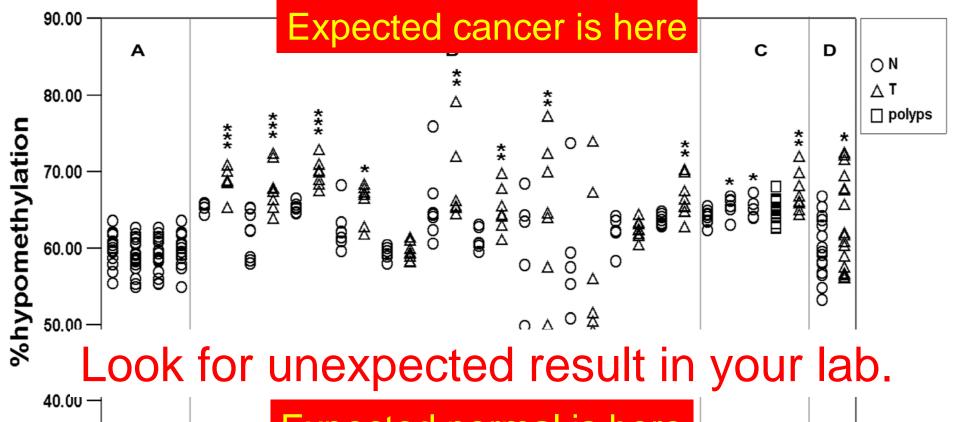


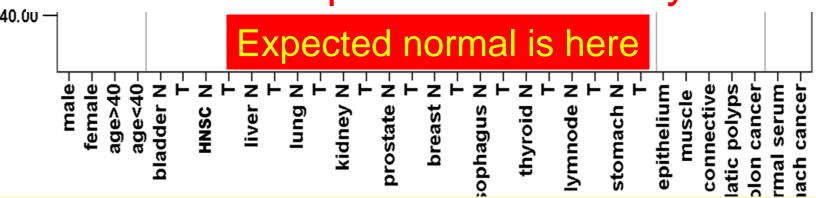


Discover of Penicillin

Sir. Alexander Fleming

"When I woke up just after dawn on September 28, 1928, I certainly didn't plan to revolutionize all medicine by discovering the world's first antibiotic, or bacteria killer," Fleming would write later, "But I guess that was exactly what I did."





Title: Distinctive pattern of LINE-1 methylation level in normal tissues and the association with

carcinogenesis

Author(s): Chalitchagorn K, Shuangshoti S, Hourpai N, et al.

Source: ONCOGENE Volume: 23 Issue: 54 Pages: 8841-8846 Published: NOV 18 2004

Times Cited: 106

ของขวัญสำหรับนักวิทย์รุ่นใหม่

- •ตาบอดคลำช้าง การทดลองด้วยเทคโนโลยี ใหม่แม้จะเป็นการศึกษาสิ่งที่เคยศึกษาแล้ว มักจะพบปรากฏการณ์ที่ไม่เคยพบมาก่อน
- •สังเกต อย่าละทิ้งสิ่งที่ไม่ได้เกิดขึ้นตามที่หวัง (เหนือความคาดหมาย)
- เข้าใจ สามารถเชื่อมโยงหาสาเหตุและ ความสำคัญของสิ่งที่สังเกตพบ

How to conduct research

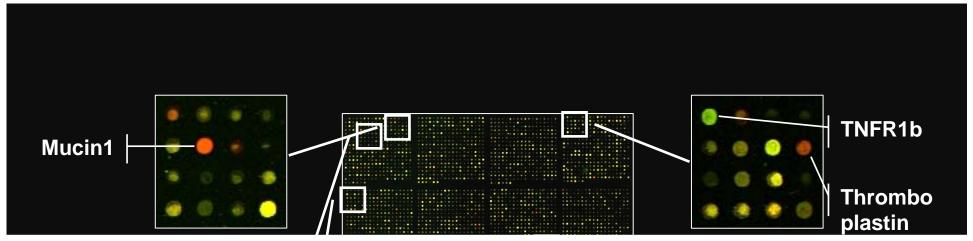
• ยืมจมูกผู้อื่นหายใจ

เพื่อ

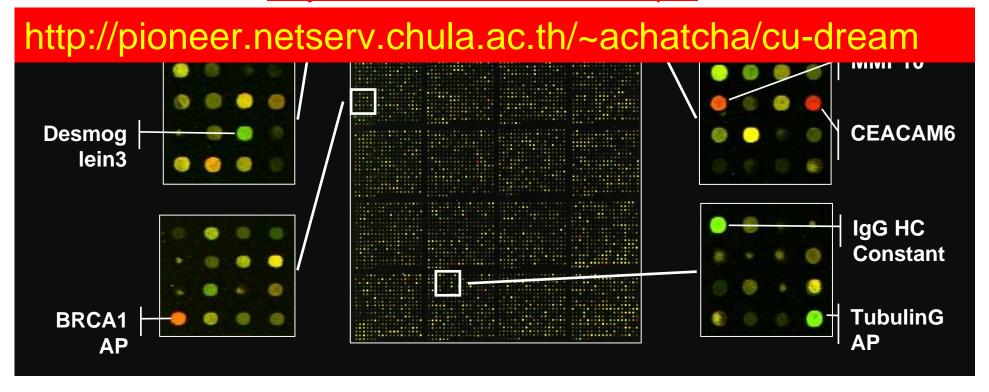
• ขนมผสมกับน้ำยา

Gingivitis vs Interferon

- -Do gingivitis increase interferon action?
- What genes are regulated by interferon in gingivitis?
- What experiments are needed?
- Several graduate students and several months or years are needed.

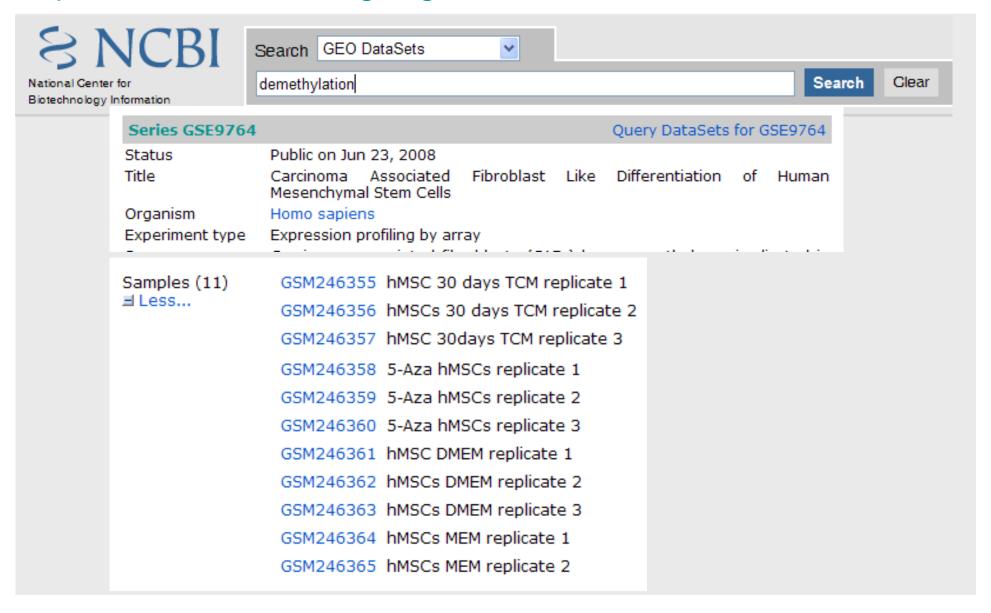


Expensive microarray expression experiments
Illumina 90,000 bahts/12 sample chip
Affymetrix 5,250 bahts/sample



CU-DREAM step 1: find GSE of interest

http://www.ncbi.nlm.nih.gov/geo/ = Gene Expression Omnibus



CU-DREAM step 2: set template & download files

1	A	В	С	D		
1	Please fill the following <u>boxes</u> .					
2		-				
3	GSE file:	GSE9764_series_matrix.txt	Series matrix file downloaded from www.ncbi.nlm.nih.gov			
4	Annotation file:	GPL570-39741.txt	Annotation file downloaded from www.ncbi.nlm.nih.gov			
5	T-test parameter (tail):	Two-tailed distribution	Parameter for ttest() in Microsoft Excel			
6	T-test parameter (type):	Paired	Parameter for ttest() in Microsoft Excel			
7	Differential expression:	Up	Direction of differential expression			
8	P-value threshold:	0.05	P-value threshold for t-test			
9						
10	Warning for paired t-test: subjects in the same row are paired.					
11		11				
12	Experimental group	Note (optional)	Control group	Note (optional)		
13	GSM246358	5-Aza hMSCs replicate 1	GSM246361	hMSC DMEM replicate 1		
14	GSM246359	5-Aza hMSCs replicate 2	GSM246362	hMSCs DMEM replicate 2		
15	GSM246360	5-Aza hMSCs replicate 3	GSM246363	hMSCs DMEM replicate 3		
16				III WARREN TO SAIT		
17						
18						
19						
20						

CU-DREAM step 3: excecute

GSE # 1
Test vs Control
T-test
Tp or Down
GSE#2
Test vs Control
Test vs Control
Up or Down
Test vs Control
Up or Down

			GSE#1
		Up	Not Up
	Up	Xxa	XXb
GSE#2	Not Up	XXc	XXd

Odd ratio > 1 or < 1 and ? significant

Gingivitis CU-DREAM Interferon Down Not down result Down 16 495 Not down 216 12,331 0.018059 P-value 1.845267 Odd Ratio 1.101765 Upper 95% CI 3.090506 Lower 95% CI d b a C SNX3 CYP2A6 RPL35 DDR1 COX4I1 RFC2 EIF4G2 RPL24 PPP1CA EIF3D HNRNPA1 HSPA6 CLPTM1 EIF3F FNTA PAX8

DDX5

OAZ1

ANAPC5

GPX1

GUCA1A

UBA7

CIB1

PMF1

Gingivitis vs Interferon

- -Do gingivitis increase interferon action? YES
- What genes are regulated by interferon in gingivitis? List provided
- What experiments are needed?
- **CU-DREAM** program and a PC
- 1 student and 10 minutes are needed.

CU-DREAM-X: excecute

GSE # X
Test vs Control
T-test
Up or Down

		GSE # X	
		Up	Not Up
Gene	list	Xxa	XXb
Not in	list	XXc	XXd

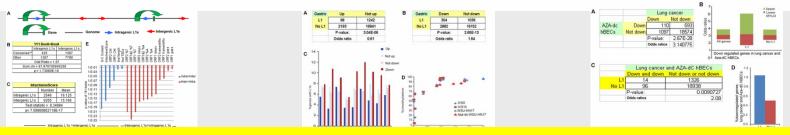
Odd ratio > 1 or < 1 and ? significant



Hypomethylation of Intragenic LINE-1 Represses Transcription in Cancer Cells through AGO2

Chatchawit Aporntewan¹⁹, Chureerat Phokaew²⁹, Jittima Piriyapongsa³⁹, Chumpol Ngamphiw³⁹, Chupong Ittiwut⁴, Sissades Tongsima³, Apiwat Mutirangura⁴*

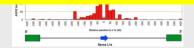
1 Department of Mathematics, Faculty of Science, Chulalongkorn University, Bangkok, Thailand, 2 Inter-Department Program of BioMedical Sciences, Faculty of Graduate School, Chulalongkorn University, Bangkok, Thailand, 3 National Center for Genetic Engineering and Biotechnology, Genome Institute, Thailand Science Park, Pathumtani, Thailand, 4 Department of Anatomy, Faculty of Medicine, Center of Excellence in Molecular Genetics of Cancer and Human Diseases, Chulalongkorn University, Bangkok, Thailand



 $402 \times 5,025 = 2,110,500$ bahts

+ sample collection and preparation





How to conduct research

• น้ำผึ้งหยดเดียว

From 1 paper Oncogene 2004, 23:8841-8846

Epigenomic patterns

Normal and cancer

Nucleic Acids Res 2008, 36:5704-5712.

Aging

Physiol Genomics 41: 194–200, 2010.

Cancer diagnosis

Clin Chim Acta 2007, 379:127-133. *Int J Gynecol Cancer* 2008, 18:711-717.

Oral Oncol 2009,

45:184-191.

Asian Pac J Cancer Prev 2007, 8:307-309. Oral Dis, 16(3):286-291. Researches in carcinogenesis (and aging)

Gene expression

Plos One

Instability & Mutation

Nucleic Acids Res 2008, **36:**3667-3675. *Molecular Cancer*

way of provention and

9(1):70.

New way of prevention and treatment of cancer (and aging)

SOME TIPS FOR BEGINNER:

Writing well:

Short and concise is always better than long and vaque.

- Verb tense: Past & present tenses.
- Introduction & Discussion: Present tense.
- Abstract, Materials/Methods and Results: Past tense, because you are describing your own work.
- When you refer to other people: you may use the past tense: "Apiwat found that this bacterium is highly sensitive to pH".
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- All tables and figures should be numbered and referenced in the text. They should also have brief captions explaining their contents.

Start drafting your pape

 Start drafting the pape vou feel most comfortable and confident with.

- Use active voice, avoid passive.
- Avoid repetition.
- Design the tables even before conducting your experiment and fill in your data when the results are obtained.
- Check all information appearing in the paper at least three times. If you have a co-author, have him/her check the information as well.
 - References, names of scientists.
 - Equations, symbols.
 - Figures caption, Tables.
 - Basis set specifications.
 - Beware! Any factual statement is the author's responsibility if it is not ascribed to somebody else through a citation. "...et al." is acceptable in the text, but not as a reference
 - Etc.,

Conclusion:

Once you have identified your favorite research areas and investigated them in detail, you can begin to control them.



Guidelines for writing a good scientific paper

Talent is a gift that many people have. It is our intention

atacticacacciataccacciccacactcattaccacciatacc

conduct the investig equipment or input files. http://biomed.md.chula.ac.th/

tnem in making the best out of it.









Peer Review: What does an editor/reviewer look for when he/she reviews a manuscript?

Is it **ORIGINAL**?

Is it **CORRECT/ACCURATE**?

Is it **INTERESTING**?

Are the data **REPRODUCIBLE**?

Are previous works properly CITED?

Will this article be cited?

Tips for writing a good scientific paper

Should be informative and "specific"

TITLE: Should be informative and "specific"

VAGUE: Chemical reactions of CNTs

SPECIFIC: Diels-Alder cycloadditions of single-wall carbon nanotubes with electron-rich dienes: a combined experimental and theoretical study

Abstract

The most important section but can be the last to be written. *Make it informative and brief.*

- Briefly state the problem or purpose of the research.
- Indicate the theoretical or experimental plan used.
- Summarize the major findings and point out major conclusions.

Introduction: In general, ovarian clear cell carcinoma (OCCC) has a history of poor response to standard platinum-based chemotherapy regimens, and advanced cases have short survival periods. Therefore, the discovery of a biomarker for the pretreatment prediction of OCCC is crucial. Loss of methylation of a retrotransposable sequence, such as long interspersed repetitive sequence 1 (LINE-1), frequently occurs in cancers, including ovarian cancer, and it has been proven to be associated with poor survival. The expressions of human endogenous retrovirus (HERV) K and E were found to be increased in tissues from patients with OCCC. Here, we propose that methylation levels of HERV are associated with treatment response and prognosis of OCCC.

Methods: Twenty-nine patients with OCCC were enrolled. Methylation levels of HERV-K, HERV-E, and LINE-1 were measured from microdissected cancer and normal ovarian tissues. The methylation levels were correlated with stage, treatment response, and prognosis.

Results: Methylation levels of HERV-K, HERV-E, and LINE-1 were decreased in tissues from patients with advanced stage cancer (P = 0.0179, P = 0.0021, and P = 0.0307, respectively). Human endogenous retrovirus K demonstrated significantly lower methylation levels in the platinum-resistant group (P = 0.0004). Patients with lower levels of methylated (hypomethylated) HERV-K had a shorter mean overall survival (P = 0.006). In advanced OCCC cases, patients with hypomethylated HERV-K had shorter mean progression-free survival (P = 0.018) and mean overall survival (P = 0.018) than did patients with higher methylation levels of HERV-K.

Conclusions: Methylation levels of HERV-K, HERV-E, and LINE-1 are decreased during OCCC multistep carcinogenesis. Moreover, HERV-K hypomethylation is a promising biomarker for predicting OCCC treatment response and prognosis.

Two to three sentences of more detailed background, comprehensible to scientists in related disciplines.

One sentence clearly stating the general problem being addressed by this particular study.

One sentence summarising the main result (with the words fliere we show or their equivalent).

Two or three sentences explaining what \
the main result reveals in direct
comparison to what was thought to be the case
previously, or how the main result adds to
previous knowledge.

One or two sentences to put the results into a more general context.

Two or three sentences to provide a broader perspective, readily comprehensible to a scientist in any discipline, may be included in the first paragraph if the editor considers that the accessibility of the paper is significantly enhanced by their inclusion. Under these circumstances, the

During cell divinion, mitoric mindles are amembled by microtubule-based motor proteins. The bipolar assanination of spindles is essential for proper settretation of chromosomes and requires plus-end-directed homotetrameric motor proteins of the widely conserved hinesin-5 (BimC) family. Hypotheses for bipolar mindle formation include the 'purh-pull mitotic murcle' model, in which hinsein-5 and opposing motor proteins act between overlapping microtubules. However, the precise roles of kinetin-5 during this process are unknown. Here we show that the vertebrate kinesin-5 Es5 drives the diding of microtubules depending on their relative orientation We found in controlled in vitre agrees that Ec5 has the remarkable capability of simultaneously moving at ~20 nm s⁻¹ towards the plus-ends of each of the two microtubules is crombule. For anti-parallel microsubuler, this results in relative rliding at =40 nm x^4 , comparable to spindle pole separation rates in vivo. Furthermore, we found that Eg5 can tether microtubule plus-ends, suspenting an additional microtubule-binding mode for Eg5. Our regults demonstrate bow members of the hinesin-5 family are likely to function in mitoria, punking apart interpolar microtubules as well as recruiting microtubules into bundles that are subsequently polarized by relative sliding. We anticipate our agray to be a starting point for more suphisticated in vitre models of mitotic pindles. For example, the individual and combined action of multiple mitoric motors could be tested, including minus-enddirected motors opposing Eg5 motility, Furthermore, Eg5 inhibition is a major target of anti-cancer drug development and a well-defined and quantitative array for motor function will be relevant for such developments.

length of the paragraph can be up to 300 words. (The above example is 190 words without the final section, and 250 words with it).



Method

Complete detail but concise.

Trick
Read as many as
possible papers with
the same approach. Then
follow the good one.

- Use specific, informative language and include precisely what you have done. This assists other scientists in reproducing your results.
- Omit unwanted information: the reader is not interested in superfluous details or asides. These waste space and raise printing costs.
- If you modified the method, say exactly what the difference of your results is from what the original method would have given, even if it is a minor one.

Introduction

• It is the place where your work is put into the proper perspective. A clearly written section builds the foundation for keeping the interest of the reader.



Writing rules: NO cookbook recipe:

Begin by answering the following Q's:

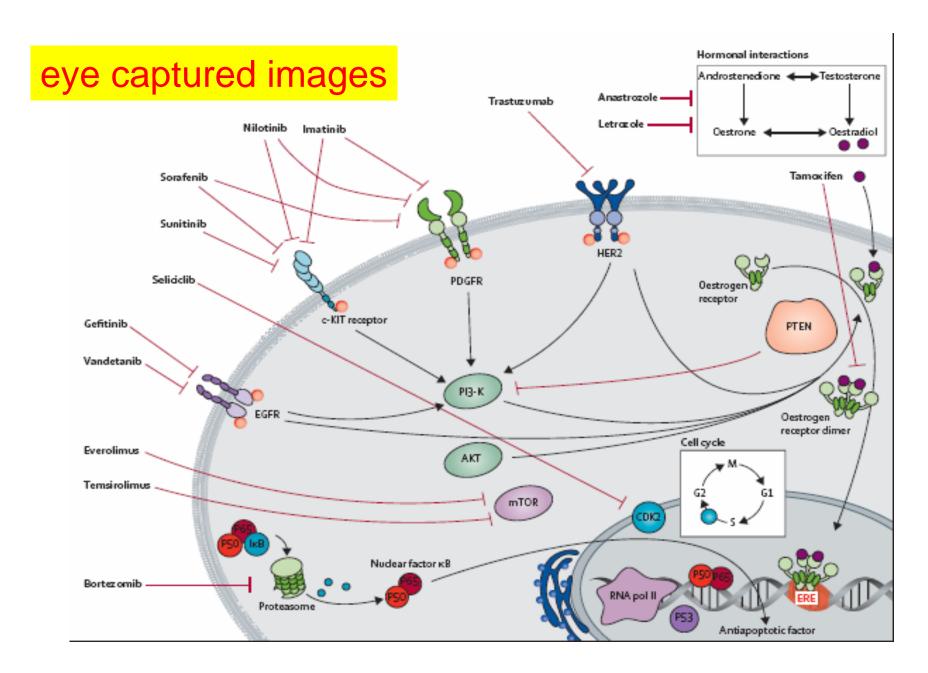
- What is new and why is your work important?
- Familiarize the reader by summarizing pertinent works in the field.
- What is already known about the system that you are investigating? Reference the most important experimental/theoretical previous work.
- How is your research significantly different from those described in the other papers?
- The contribution that your paper will have to the advancement of science and technology in general.
- What are the major objectives of your paper?

For those who are new.

- 1. Gather information from literatures.
- 2. Short note sentences to be cited.
- 3. Grouping information together with planning sequence of paragraphs.
- 4. Structure each paragraph with topic sentence, detail and conclusion.

Illustrations (from hand out)

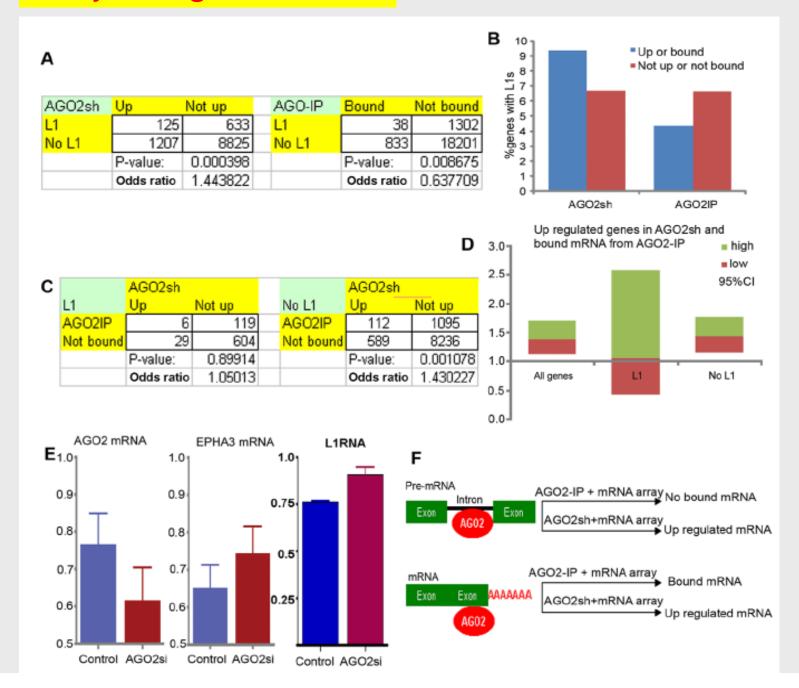
"One picture is indeed better than a thousand words".



Manchana T, Ittiwut C, Mutirangura A, Kavanagh JJ: Targeted therapies for rare gynaecological cancers. Lancet Oncology 2010, 11(7):685-693.

storytelling illustration

PLoS One 2011, 6(3):e17934.



Result

- Summarize and illustrate your findings.
- Integrate quantitative data with the text.
- Similar to you are telling interesting story or making the story interesting.
- Thus result should include a bit of introduction, method and discussion.
- More importantly there should be interesting scientific reason connecting between each section. So it should look as if your first experimental result lead to the second and the second lead to third and fourth, respectively.

Discussion (from Hand-out)

- Interpret your findings and support your conclusions with solid evidence (DO NOT speculate).
- Refer to your data, citing tables and figures where necessary; use these materials as evidence to support your major arguments.

Discussion

- Analysis (Interpret your findings and support your conclusions with solid evidence (DO NOT speculate).
- Comparison (with others)
- Advertise (How do your results fit into the bigger picture?)
- Correction and declare limitation (only if needed or reviewers request.)
- Discussion แปลว่าวิจารณ์ไม่ใช่จับผิด

Remarks:

- Are your findings consistent with those of other researchers?
- How do your results fit into the bigger picture? (defined in the Introduction).
- Do not present every conceivable explanation.
- Recognize the importance of "negative" results.
- Cases that do not conform to the expected pattern might represent something "breaking a new ground".
- No section or subsection of the paper should be more than five pages long, around a total of 15 pages. You need to help the readers find the material they want.
- Papers have to end up with a section called "Conclusions".

Papers have to end up with a section called "Conclusions"

Conclusion 3'Rs

Restate: your findings more concisely

Recap: give a summary of the main points of the discussion

Recommend & benefit: e.g.,

The results derived in the present study suggest that the KU-method yields: an accurate and practical model for exploring the reaction mechanisms of nanostructured zeolite catalysts.

Supplements

Reviewers do not read it.

The more you have the better your study looks

Letter to editor

Very important.

The more editor likes your work, the better chance to get reasonable reviewers.



Peer Review: What does an editor/reviewer look for when he/she reviews a manuscript?

Is it **ORIGINAL**?

Is it **CORRECT/ACCURATE**?

Is it **INTERESTING**?

Are the data **REPRODUCIBLE**?

Are previous works properly CITED?

Will this article be cited?

Suggest reviewers

Suggest who you met.

If you do not know anyone, don't try

Keywords

Acknowledgements

Editing

- Grammar and spelling
- Concise (giving much information in few words), Thesaurus
- Be patient.
- Ask your English spoken friends for help.
- Try professional

SOME TIPS FOR BEGINNER:

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Past & present tenses.

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- Start drafting the paper from the section that you feel most comfortable and confident with.
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Talent is a gift that many people have. It is our intention to identify such talent in young

people and assist them in making the best out of it.





Guidelines for writing a good scientific paper





Supports by

- "Research Chair Grant 2011, National Science and Technology Development Agency (NSTDA),"
- "Senior Research Scholar TRF"
- " Four Seasons Hotel Bangkok's 4th Cancer Care charity fun run in coordination with the Thai Red Cross Society."

Center of Excellence in Molecular Genetics of Cancer and Human DiseasesDepartment of Anatomy, Faculty of Medicine Chulalongkorn University



กิตติกรรมประกาศ

ขอขอบคุณครูพันธุศาสตร์ Prof. David H Ledbetter, Prof. Arthur L. Beaudet และ ผศ. พญ. อรศรี รมยะนันท์ ขอขอบคุณความร่วมมือที่ดีเยี่ยมจากสมาชิกผู้วิจัยที่เป็นอาจารย์ แพทย์ ้นักวิทยาศาสตร์ ส่วนหนึ่งได้แก่ รศ.นพ. ภาคภูมิ สุปิยพันธุ์ รศ.พญ. เสาวณีย์ เย็นฤดี รศ.นพ. นรินทร์ วรวุฒิ ผศ.พญ. กาญจนา โชติเลอศักดิ์ รศ.นพ. วีรชัย ศิริกาญจนะรงค์ รศ.นพ. สมชัย นิรุตติศาสน์ รศ.นพ. ดำรง ตรีสุโกศล รศ.นพ. ประเสริฐ ตรีวิจิตรศิลป์ รศ.พญ. สุรางค์ ตรีรัตน ชาติ ผศ.นพ. วิชัย เติมรุ่งเรืองเลิศ ศ.นพ. นเรศร สุขเจริญ รศ.นพ. ประเสริฐ เลิศสงวนสินชัย รศ.นพ. สังคม จงพิพัฒน์วณิชย์ ศ.นพ. วรศักดิ์ โชติเลอศักดิ์ อ.พญ. รัศมน กัลยาศิริ อ.ดร. ทพญ อรนาฎ มาตังคสมบัติ รศ.นพ. พิสิฐ ตั้งกิจวานิชย์ ศ.นพ. ยง ภู่วรวรรณ ผศ.นพ. ชล ้เกียรติ ขอประเสริฐ อ.ดร.พญ. จงกลนี วงศ์ปิยะบวร น.ส. สายรุ้ง ศักดิกุล นายประกาศิต รัตน ์ ตันหยง อ.นพ. สรภพ เกียรติพงศ์สาน รศ.นพ. อิศรางค์ นุชประยูร น.ส. นุสรา หัวไผ่ Ms Petra Hirsch, Dr. J. Silvio Gutkind, น.ส. ชุติภา พวงไพโรจน์ น.ส. ธัชวรรณ ธนาศุภวัฒน์ น.ส. โชติกา สุญาณเศรษฐกร นาย ศุภกิจโขวุฒิธรรม ศ.ทพญ. กอบกาญจน์ ทองประสม ดร ์แสงสูรย์ เจริญวิไลศิริ รศ.ดร.พญ. ณัฏฐิยา หิรัญกาญจน์ ศ.นพ. ชนพ ช่วงโชติและ ผศ.ดร. นพ. วิโรจน์ ศรีอุฬารพงศ์ ให้คำปรึกษาทางคลินิกและร่วมทำวิจัย ขอขอบคุณความร่วมมือที่ดีเยี่ยมจากสมาชิกนิสิตระดับบัณฑิตศึกษา บัณฑิตที่จบการศึกษา ส่วนหนึ่งได้แก่ ดร. วิชัย พรธนเกษม น.ส. วันเพ็ญ พลเยี่ยม และ ดร. นริศร คงรัตนโชค ศึกษา หมู่เมททิลของ EDSBs. น.ส. พิชญาณีน ชลิตชากร พญ. ฐิติวรรณ ลมดี อ.ทพญ. ดร. เกศ กัญญา สัพพะเลข พญ. จีริชุดา ปัทมดิลก และ น.ส. จุรีรัตน์ โพธิ์แก้ว ศึกษาปริมาณและ ลักษณะการกระจายตัวของหมู่เมททิลที่ไลน์ 1 ดร. เกรียงศักดิ์ ฤชุศาศวัต น.ส. รุ่งทิวา วัฒนวราภรณ์ ดร. ปฐมวดี ญาณทัศนีย์จิต ดร. จิรนันท์ วราชิต และ อ.ดร.ทพ. นครินทร์ กิต ้กำธร ศึกษาหมู่เมททิลของยืน ดร.ชูพงศ์ อิทธิวุฒิ และ ดร. รุ่งนภา หิรัญสถิต ศึกษาพันธุ ์ศาสตร์ของประชากร ผศ.ดร.พญ. มนตกานต์ ตันสถิตย์ ศึกษาพันธุศาสตร์ของเซลล์ ดร. วิชัย พรธนเกษม ดร. นริศร คงรัตนโชค ดร. รุ่งนภา หิรัญสถิต และ อ.พญ. จันทร์ทรา ตนันท์ยุทธ ส์ สืดแดนเรียโพธ เหด้าดนด