





Resting-State fMRI: Applications

Chao-Gan YAN, Ph.D.
严超赣
ycg.yan@gmail.com
http://rfmri.org
Institute of Psychology, Chinese Academy of Science



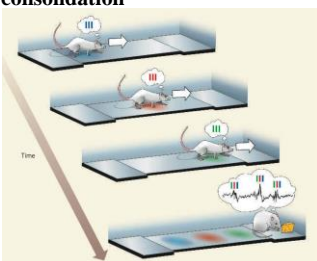
Outline

- ➔ • Applications to Cognitive Science
- Applications to Brain Disorders




Applications to Cognitive Science

Off-line spontaneous brain activity and memory consolidation



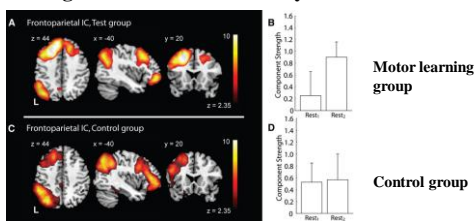
Reverse replay of
behavioral sequences in
hippocampal place cells
during the awake state.

Foster and Wilson, 2006. Nature; Colgin and Moser, 2006. Nature




Applications to Cognitive Science

Resting-state fMRI and memory consolidation



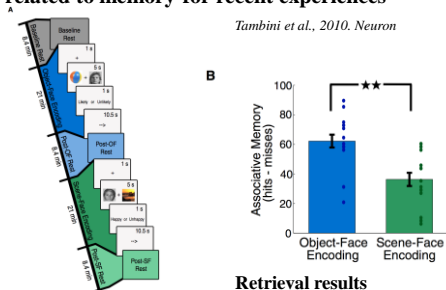
Motor learning but not motor performance modulated
subsequent frontal-parietal resting-state network

Albert et al., 2009. Curr Biol




Applications to Cognitive Science

Enhanced brain correlations during rest are related to memory for recent experiences

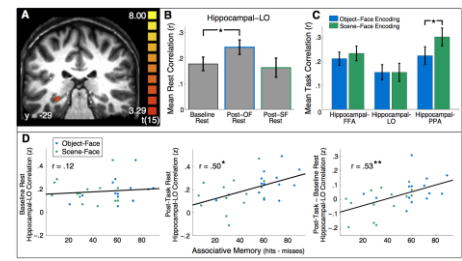


Retrieval results


Tambini et al., 2010. Neuron



Applications to Cognitive Science



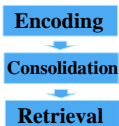
Tambini et al., 2010. Neuron



Applications to Cognitive Science

Episodic memory and Alzheimer disease:

- Subtle change of episodic memory is the earliest cognitive deficit in AD (Schwindt & Black, 1999, *NeuroImage*)
- Consistently decreased activation in AD patients during both encoding and retrieval stages of episodic memory (Schwindt & Black, 2009, *NeuroImage*)
- Mechanism of episodic memory consolidation after encoding?



Applications to Cognitive Science

Resting-state fMRI, memory consolidation, APOE ε4

- Question-1: Resting-state fMRI, hippocampus activity, and APOE ε4 healthy carriers?
- Question-2: Spontaneous activity modulation by episodic memory task in the brain regions for encoding?



Applications to Cognitive Science

Participants:

- 917 students
2/2 2/3 2/4 3/3 3/4 4/4
1 52 13 799 51 1
- Two groups of healthy APOE carriers: ε4/3 vs. ε2/3, n = 20 vs 19, (half males), 18 – 23 yrs from BNU, match for IQ and education

Wang, , Yan et al., 2012. *PLoS ONE*



Applications to Cognitive Science

Design

- Scanning sessions (3T Siemens):
- | | | |
|------|---|--------|
| ➡ S1 | Resting-state (Rest1) | 8 min |
| S2 | Pictures (indoor or outdoor) (encoding) | 5 min |
| S3 | 3D structure | 8 min |
| ➡ S4 | Resting-state (Rest2) | 8 min |
| S5 | Retrieval: old or new (2 runs) | 10 min |
- Rest2/Rest1: spontaneous brain activity modulation by task

Wang, , Yan et al., 2012. *PLoS ONE*



Applications to Cognitive Science

Behavior result: retrieval performance

	d' mean	SD	t	p
2/3	2.46	0.68	-0.387	0.7
3/4	2.54	0.63		

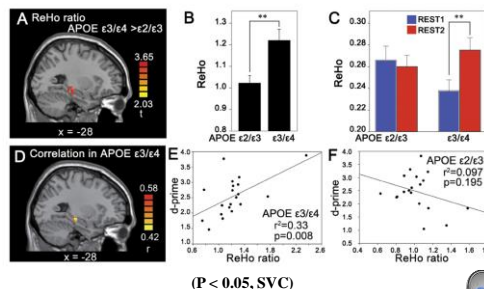
No significant difference between the two groups

Wang, , Yan et al., 2012. *PLoS ONE*



Applications to Cognitive Science

Hippocampus



($P < 0.05$, SVC)

Wang, , Yan et al., 2012. *PLoS ONE*



Outline

- Applications to Cognitive Science

- ➔ • Applications to Brain Disorders



Applications to Brain Disorders

- Alzheimer's Dementia (AD)
- Depression
- Autism Spectrum Disorder (ASD)
- ...

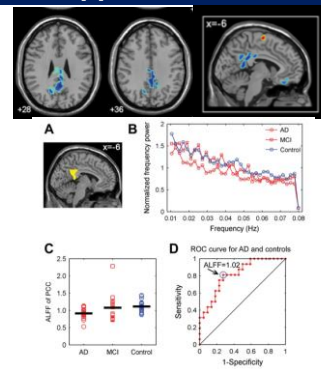


Applications to Brain Disorders

ALFF in AD



Applications to Brain Disorders

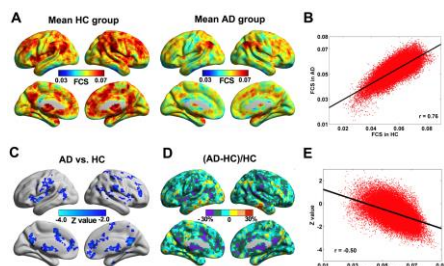


Wang*, Yan* et al., 2011,
Hum Brain Mapp

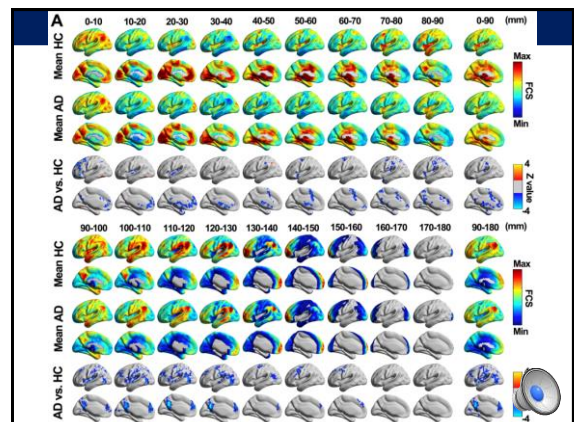


Applications to Brain Disorders

Degree centrality in AD



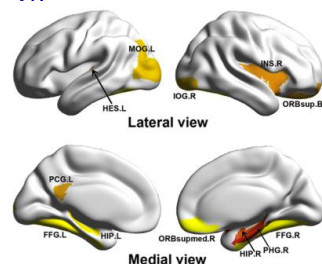
Dai, Yan et al., 2014. Cerebral Cortex.



Applications to Brain Disorders

Classification

- Multi-modal characterization
- Features
 - ALFF
 - ReHo
 - Region
 - Gray matter
- Accuracy: 89.47%

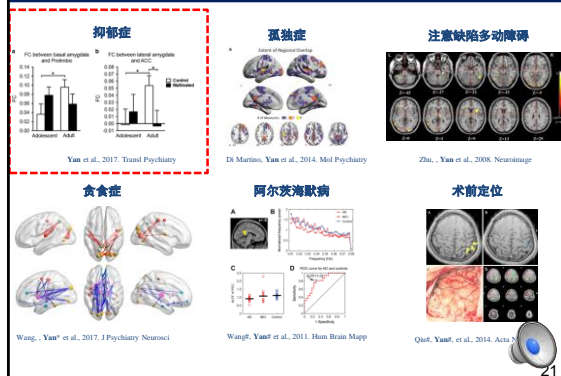


Dai[#], Yan[#] et al., 2012, NeuroImage

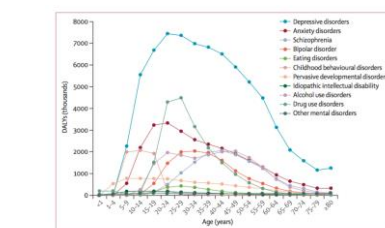
Applications to Brain Disorders

- Alzheimer's Dementia (AD)
- Depression
- Autism Spectrum Disorder (ASD)
- ...

广泛应用于脑疾病中

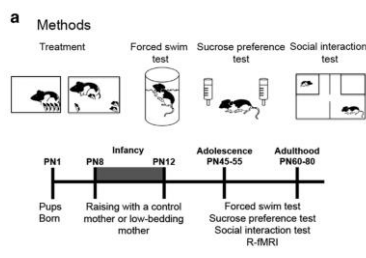


MDD疾病负担重，诊断缺乏生物学客观标准



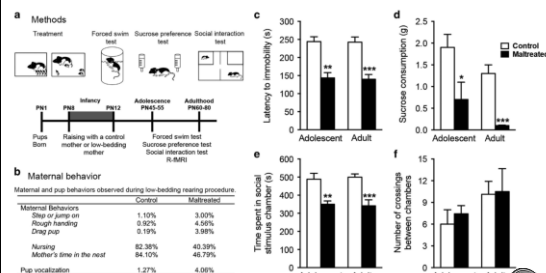
抑郁症（重性抑郁障碍，MDD）疾病负担居于首位

Depressive-like behavior in rat model of caregiver maltreatment



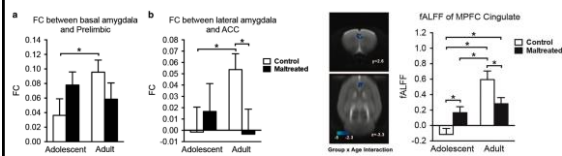
Yan et al., 2017, Transl Psychiatry

Depressive-like behavior in rat model of caregiver maltreatment



Yan et al., 2017, Transl Psychiatry

Depressive-like behavior in rat model of caregiver maltreatment

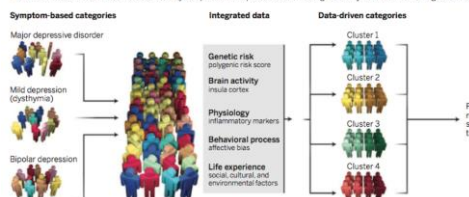


Yan et al., 2017. Transl Psychiatry

The R-fMRI Maps Project Consortium for Depression and Bipolar Disorders

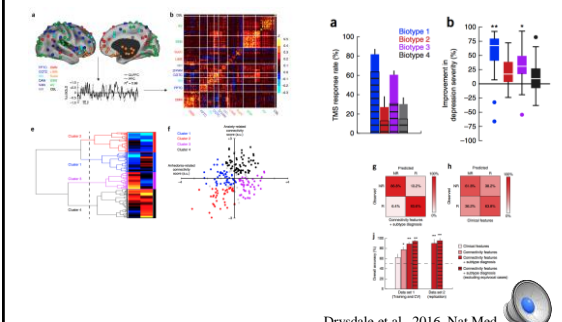
Deconstructed, parsed, and diagnosed.

A hypothetical example illustrates how precision medicine might deconstruct traditional symptom-based categories. Patients with a range of mood disorders are studied across several analytical platforms to parse current heterogeneous syndromes into homogeneous clusters.



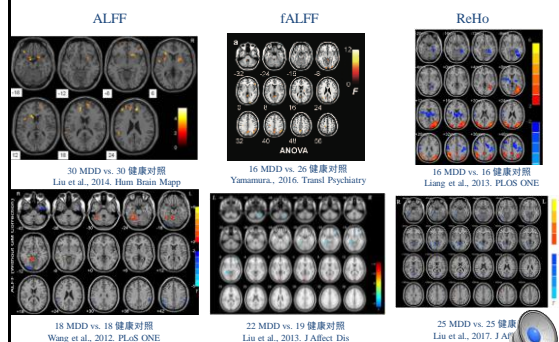
Insel and Cuthbert 2015. Science

Biotypes in Depression



Drysdale et al., 2016. Nat Med

MDD静息态功能磁共振研究



28

小样本研究遇到的问题

ANALYSIS

Power failure: why small sample size undermines the reliability of neuroscience

Kohonen V, Ballester C, John P A, Savadkou S, Cline M, Wang B, Ryan A, Neeb R, Jendryak J, Ermiu S, J. Robinson and Marcus R. Munoz

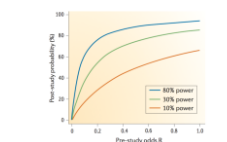
Button et al., 2013. Nat Rev Neurosci

ANALYSIS

Scanning the horizon: towards transparent and reproducible neuroimaging research

Russell A, Poldrack R, Chen L, Baker J, John D, Durrant J, Kogut J, Gorgolewski J, ...

Poldrack et al., 2017. Nat Rev Neurosci



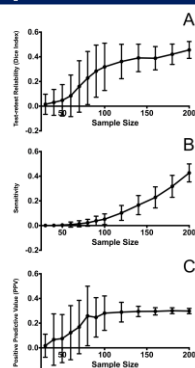
- 统计力不足时，发现的结果为真的可能性很小
- 神经科学领域统计力约为8%-31%

2015年，fMRI研究每样本的中位数仅为19

29

Sample Size Matters

Randomly draw k subjects from the "SWU 4" site in the CORR dataset, which has two sessions of 116 males and 105 females



Chen, Lu, Yan, 2017. Human Brain Mapping

30

抑郁症静息态功能磁共振多中心大数据研究



31

研究方案



获得国内24家医院33位精神科专家的支持



2017年3月25日，抑郁症静息态功能磁共振多中心数据荟萃分析会议于中科院心理所成功召开

赵靖平 中南大学湘雅二医院
谢光荣 中南大学湘雅二医院
刘哲宁 中南大学湘雅二医院
王传联 安定医院
陈伟 浙江大学邵逸夫医院
司天梅 北京大学第六医院
蒙华庆 重庆医科大学
戚玉峰 杭州师范大学
傅小兰 中科院心理所
刘勋 中科院心理所



33

抑郁症静息态功能磁共振多中心数据荟萃分析(REST-meta-MDD)计划参与单位



北京大学第六医院、北京安定医院、中南大学湘雅二医院、四川大学华西医院、东南大学附属中大医院、浙江大学医学院附属第一医院与附属邵逸夫医院、南京脑科医院、苏州广济医院、暨南大学附属第一医院、昆明医学院第一附属医院、上海交通大学医学院附属精神卫生中心、首都医科大学附属北京安定医院、四川大学华西医院、苏州市广济医院、西安市中心医院、西南大学心理学院、浙江大学医学院附属第一医院、浙江大学医学院附属邵逸夫医院、中国医科大学第一附属医院、中国医科大学、安徽医科大学、山西医科大学、昆明医科大学、西安交通大学、西南大学、复旦大学、杭州师范大学.....



已成功累积1300例MDD与1128例正常对照数据，为全球最大的抑郁症静息态功能磁共振数据库

34

REST-meta-MDD

表1. REST-meta-MDD 计划参与研究单位以及数据构成情况

编号	参与研究单位	样本量	
		MDD	NC
1	安徽医科大学	51	36
2	北京大学第六医院	74	74
3	重庆医科大学附属第一医院	111	79
4	东南大学附属中大医院	139	113
5	暨南大学第一附属医院	50	50
6	昆明医学院第一附属医院	32	31
7	山西医科大学第一医院	50	33
8	上海交通大学医学院附属精神卫生中心	28	26
9	首都医科大学附属北京安定医院	86	70
10	四川大学华西医院	63	61
11	苏州市广济医院	30	30
12	西安市中心医院	25	17
13	西南大学心理学院	282	251
14	浙江大学医学院附属第一医院	21	20
15	浙江大学医学院附属邵逸夫医院	38	49
16	中国医科大学第一附属医院	75	75
17	中南大学湘雅二医院	145	113

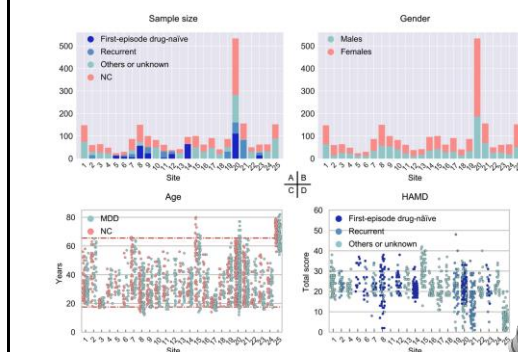
注：MDD：重度抑郁障碍；NC：健康对照

Yan et al., in prep.



35

REST-meta-MDD



Yan et al., in prep.



36

异常脑区与抑郁心理加工机制的关联

反刍思维是指对
负性生活事件及其
意义、原因、
后果的反复思考



静息态

悲伤事件
回忆

反刍思
维状态

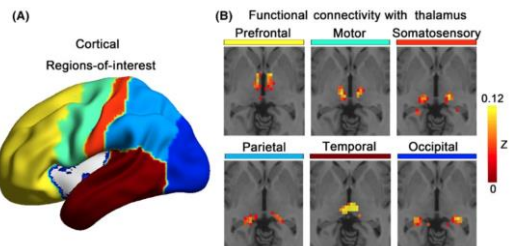
快乐事
件回忆



Nolen-Hoeksema et al., 2008. Perspect Psychol Sci; Christoff et al., 2016. Nature Rev Neurosci

43

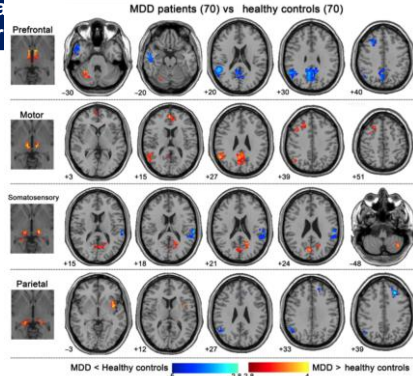
Aberrant intrinsic functional connectivity in thalamo-cortical networks in major depressive disorder



Kong, Yan*, Si*, 2018. CNS Neurosci Ther

44

Aberrant intrinsic functional connectivity in thalamo-cortical networks in major depressive disorder



Kong, Yan*, Si*, 2018. CNS Neurosci Ther

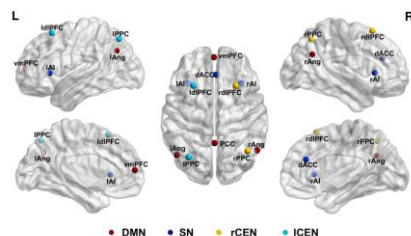
45

Applications to Brain Disorders

- Alzheimer's Dementia (AD)
- Depression
- Autism Spectrum Disorder (ASD)
- ...



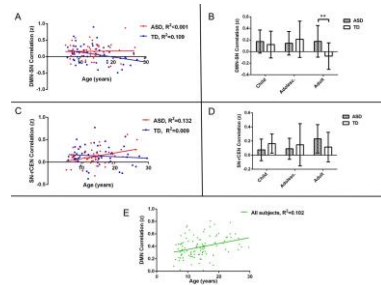
Altered Developmental Trajectories in Intrinsic Function between Default, Salience, and Executive Networks in High-Functioning Autism



Yang, Yan*, in prep.

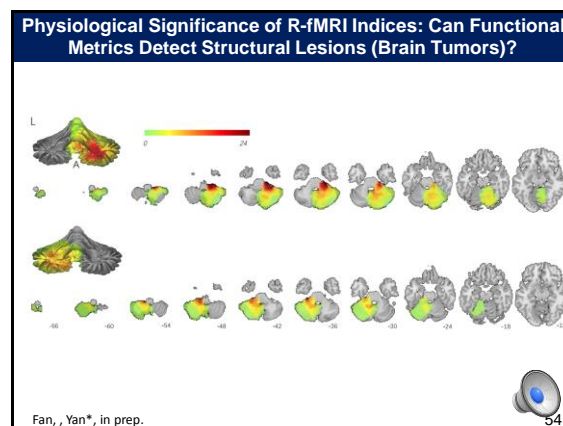
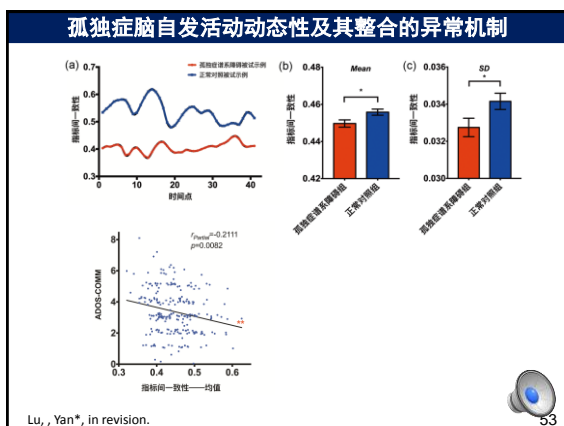
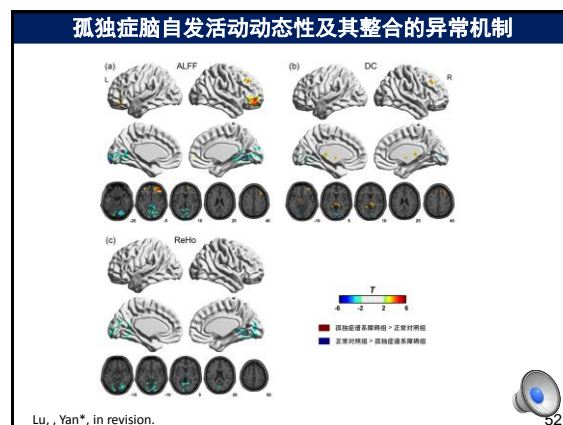
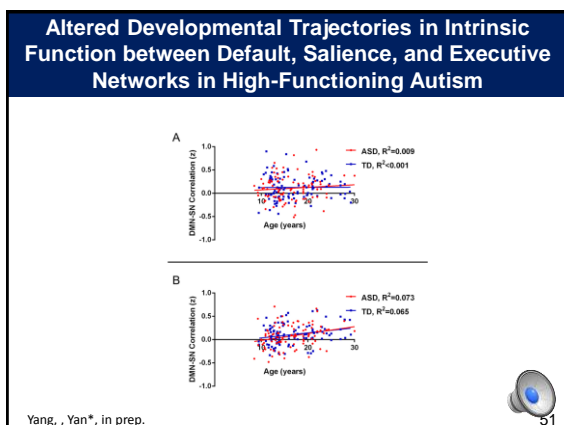
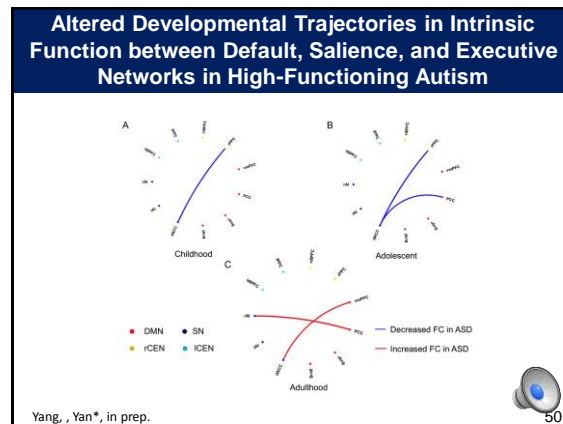
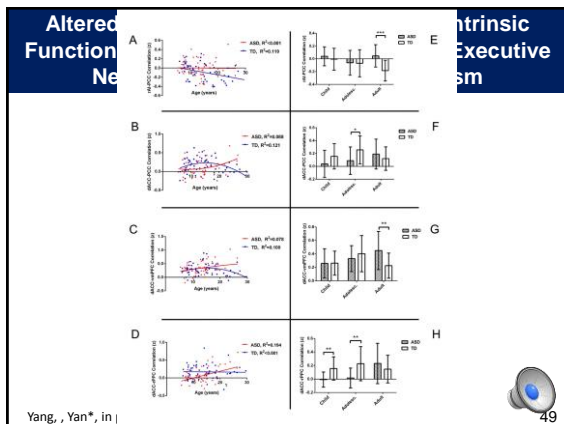
47

Altered Developmental Trajectories in Intrinsic Function between Default, Salience, and Executive Networks in High-Functioning Autism

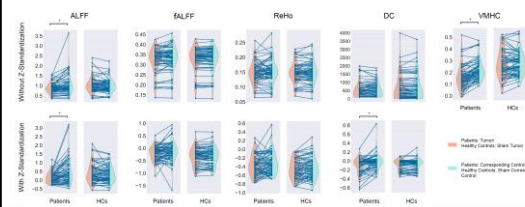


Yang, Yan*, in prep.

48



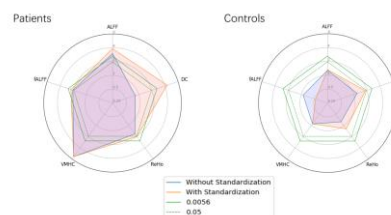
Physiological Significance of R-fMRI Indices: Can Functional Metrics Detect Structural Lesions (Brain Tumors)?



Fan, Yan*, in prep.

55

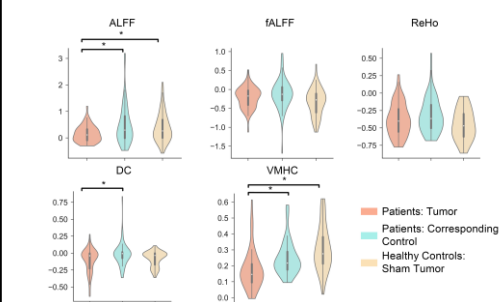
Physiological Significance of R-fMRI Indices: Can Functional Metrics Detect Structural Lesions (Brain Tumors)?



Fan, Yan*, in prep.

56

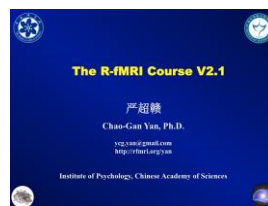
Physiological Significance of R-fMRI Indices: Can Functional Metrics Detect Structural Lesions (Brain Tumors)?



Fan, Yan*, in prep.

57

Further Help


<http://rfmri.org/Course>

<http://rfmri.org/wiki>


The R-fMRI Journal Club



Official Account: RfMRI Lab

58

Preprints of the R-fMRI Network



Preprints of the R-fMRI Network (PRN) is a preprint, open-access, free-submission, open-discussion, community funded Preprints of R-fMRI related research. The goal of PRN is to supplement the peer reviewed journal publication system – by more rapidly communicating the latest research achievements across the global.

F1000Research

F1000Research 2018, 8(12):e61461, 21 Aug 2019



SOFTWARE TOOL ARTICLE

REVIEWED PRN: a preprint service for catalyzing R-fMRI and neuroscience related studies [v2; ref status: indexed, <http://f1000r.es/5qy>]

Chao-gan Yan^{1,4}, Qingyang Li⁴, Lei Gao^{4,5}¹The Nathan Kline Institute for Psychiatric Research, Orangeburg, NY, USA²Institute of Psychology, Chinese Academy of Sciences, 16 Lincui Rd, Chaoyang District, Beijing, 100101, China³Department of Child and Adolescent Psychiatry, New York University Langone Medical Center, New York, NY, USA⁴National Office of PRN, the R-fMRI Network, Inc., New York, NY, USA⁵Department of Radiology, the First Affiliated Hospital of Nanchang University, Nanchang, China

59

数据分析与深度培训



静态功能磁共振成像深度数据分析

功能磁共振成像越来越成为一种主流的科技手段，然而功能磁共振的数据分析却是一项具有高度挑战性的工作。海量的原始数据，繁多的分析步骤，复杂的分析方法都让研究者们无所适从。恰当的分析方法可以从海量的数据中挖掘出富有创新性的结果，而不恰当的分析则可能让精心收集的数数据黯然失色。深度大脑公司联合中国科学院 The R-fMRI Lab 的专业脑功能磁共振研究团队，推出一站式功能磁共振数据分析解决方案，助您从容应对功能磁共振数据带来的挑战。

<http://deepbrain.com>


静态功能磁共振成像数据处理深度培训

从您见到这条消息开始，您便将有与中国科学院 The R-fMRI Lab 的静态功能磁共振专家团队合作探索大脑的奥秘！深度跟组特训期间，您将会亲身体验：

- 数据处理 专家指导下高效学习静态功能磁共振成像数据处理
- 思路设计 与国际知名专家讨论研究思路
- 论文撰写 系统的 SCI 论文写作训练



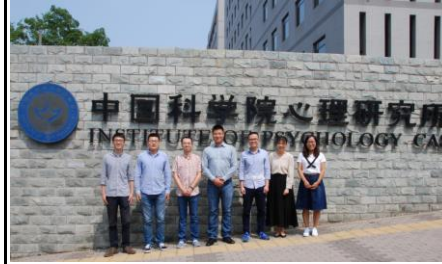
The R-fMRI Lab



WeChat Official Account: RFMRILab



Acknowledgments



Chinese Academy of
Sciences
Xi-Nian Zuo
Hangzhou Normal
University
Yu-Feng Zang
NYU Child Study
Center
F. Xavier Castellanos
Child Mind Institute
Michael P. Milham

- Funding**
- National Natural Science Foundation of China
 - National Key R&D Program of China
 - Chinese Academy of Sciences



Thanks for your attention!

